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#### (54) Title: 5' ESTs FOR SECRETED PROTEINS EXPRESSED IN BRAIN

#### (57) Abstract

The sequences of 5' ESTs derived from mRNAs encoding secreted proteins are disclosed. The 5' ESTs may be to obtain cDNAs and genomic DNAs corresponding to the 5' ESTs. The 5' ESTs may also be used in diagnostic, forensic, gene therapy, and chromosome mapping procedures. Upstream regulatory sequences may also be obtained using the 5' ESTs. The 5' ESTs may also be used to design expression vectors and secretion vectors.

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#### 5' ESTs FOR SECRETED PROTEINS EXPRESSED IN BRAIN

#### Background of the Invention

The estimated 50,000-100,000 genes scattered along the human chromosomes offer tremendous promise for the understanding, diagnosis, and treatment of human diseases. In addition, probes capable of specifically hybridizing to loci distributed throughout the human genome find applications in the construction of high resolution chromosome maps and in the identification of individuals.

In the past, the characterization of even a single human gene was a painstaking process, requiring years of effort. Recent developments in the areas of cloning vectors, DNA sequencing, and computer technology have merged to greatly accelerate the rate at which human genes can be isolated, sequenced, mapped, and characterized. Cloning vectors such as yeast artificial chromosomes (YACs) and bacterial artificial chromosomes (BACs) are able to accept DNA inserts ranging from 300 to 1000 kilobases (kb) or 100–400 kb in length respectively, thereby facilitating the manipulation and ordering of DNA sequences distributed over great distances on the human chromosomes. Automated DNA sequencing machines permit the rapid sequencing of human genes. Bioinformatics software enables the comparison of nucleic acid and protein sequences, thereby assisting in the characterization of human gene products.

Currently, two different approaches are being pursued for identifying and characterizing the genes distributed along the human genome. In one approach, large fragments of genomic DNA are isolated, cloned, and sequenced. Potential open reading frames in these genomic sequences are identified using bioinformatics software. However, this approach entails sequencing large stretches of human DNA which do not encode proteins in order to find the protein encoding sequences scattered throughout the genome. In addition to requiring extensive sequencing, the bioinformatics software may mischaracterize the genomic sequences obtained. Thus, the software may produce false positives in which noncoding DNA is mischaracterized as coding DNA or false negatives in which coding DNA is mischaracterized as non-coding DNA.

An alternative approach takes a more direct route to identifying and characterizing human genes. In this approach, complementary DNAs (cDNAs) are synthesized from isolated messenger RNAs (mRNAs) which encode human proteins. Using this approach,

sequencing is only performed on DNA which is derived from protein coding portions of the genome. Often, only short stretches of the cDNAs are sequenced to obtain sequences called expressed sequence tags (ESTs). The ESTs may then be used to isolate or purify extended cDNAs which include sequences adjacent to the EST sequences. The extended cDNAs may contain all of the sequence of the EST which was used to obtain them or only a portion of the sequence of the EST which was used to obtain them. In addition, the extended cDNAs may contain the full coding sequence of the gene from which the EST was derived or, alternatively, the extended cDNAs may include portions of the coding sequence of the gene from which the EST was derived. It will be appreciated that there may be several extended cDNAs which include the EST sequence as a result of alternate splicing or the activity of alternative promoters.

In the past, these short EST sequences were often obtained from oligo-dT primed cDNA libraries. Accordingly, they mainly corresponded to the 3' untranslated region of the mRNA. In part, the prevalence of EST sequences derived from the 3' end of the mRNA is a result of the fact that typical techniques for obtaining cDNAs are not well suited for isolating cDNA sequences derived from the 5' ends of mRNAs. (Adams *et al.*, *Nature* 377:3-174, 1996; Hillier *et al.*, *Genome Res.* 6:807-828, 1996).

In addition, in those reported instances where longer cDNA sequences have been obtained, the reported sequences typically correspond to coding sequences and do not include the full 5' untranslated region of the mRNA from which the cDNA is derived. Such incomplete sequences may not include the first exon of the mRNA, particularly in situations where the first exon is short. Furthermore, they may not include some exons, often short ones, which are located upstream of splicing sites. Thus, there is a need to obtain sequences derived from the 5' ends of mRNAs.

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While many sequences derived from human chromosomes have practical applications, approaches based on the identification and characterization of those chromosomal sequences which encode a protein product are particularly relevant to diagnostic and therapeutic uses. Of the 50,000-100,000 protein coding genes, those genes encoding proteins which are secreted from the cell in which they are synthesized, as well as the secreted proteins themselves, are particularly valuable as potential therapeutic agents. Such proteins are often

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involved in cell to cell communication and may be responsible for producing a clinically relevant response in their target cells.

In fact, several secretory proteins, including tissue plasminogen activator, G-CSF, GM-CSF, erythropoietin, human growth hormone, insulin, interferon-α, interferon-β, interferon-γ, and interleukin-2, are currently in clinical use. These proteins are used to treat a wide range of conditions, including acute myocardial infarction, acute ischemic stroke, anemia, diabetes, growth hormone deficiency, hepatitis, kidney carcinoma, chemotherapy induced neutropenia and multiple sclerosis. For these reasons, extended cDNAs encoding secreted proteins or portions thereof represent a particularly valuable source of therapeutic agents. Thus, there is a need for the identification and characterization of secreted proteins and the nucleic acids encoding them.

In addition to being therapeutically useful themselves, secretory proteins include short peptides, called signal peptides, at their amino termini which direct their secretion. These signal peptides are encoded by the signal sequences located at the 5' ends of the coding sequences of genes encoding secreted proteins. Because these signal peptides will direct the extracellular secretion of any protein to which they are operably linked, the signal sequences may be exploited to direct the efficient secretion of any protein by operably linking the signal sequences to a gene encoding the protein for which secretion is desired. In addition, portions of signal sequences may also be used to direct the intracellular import of a peptide or protein of interest. This may prove beneficial in gene therapy strategies in which it is desired to deliver a particular gene product to cells other than the cell in which it is produced. Signal sequences encoding signal peptides also find application in simplifying protein purification techniques. In such applications, the extracellular secretion of the desired protein greatly facilitates purification by reducing the number of undesired proteins from which the desired protein must be selected. Thus, there exists a need to identify and characterize the 5' portions of the genes for secretory proteins which encode signal peptides.

Public information on the number of human genes for which the promoters and upstream regulatory regions have been identified and characterized is quite limited. In part, this may be due to the difficulty of isolating such regulatory sequences. Upstream regulatory sequences such as transcription factor binding sites are typically too short to be utilized as probes for isolating promoters from human genomic libraries. Recently, some approaches

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have been developed to isolate human promoters. One of them consists of making a CpG island library (Cross, et al., Nature Genetics 6: 236-244, 1994). The second consists of isolating human genomic DNA sequences containing SpeI binding sites by the use of SpeI binding protein. (Mortlock et al., Genome Res. 6:327-335, 1996). Both of these approaches have their limits due to a lack of specificity or of comprehensiveness.

The present 5' ESTs may be used to efficiently identify and isolate upstream regulatory regions which control the location, developmental stage, rate, and quantity of protein synthesis, as well as the stability of the mRNA. (Theil, *BioFactors* 4:87-93, 1993). Once identified and characterized, these regulatory regions may be utilized in gene therapy or protein purification schemes to obtain the desired amount and locations of protein synthesis or to inhibit, reduce, or prevent the synthesis of undesirable gene products.

In addition, ESTs containing the 5' ends of secretory protein genes may include sequences useful as probes for chromosome mapping and the identification of individuals. Thus, there is a need to identify and characterize the sequences upstream of the 5' coding sequences of genes encoding secretory proteins.

#### Summary of the Invention

The present invention relates to purified, isolated, or recombinant ESTs which include sequences derived from the authentic 5' ends of their corresponding mRNAs. The term "corresponding mRNA" refers to the mRNA which was the template for the cDNA synthesis which produced the 5' EST. These sequences will be referred to hereinafter as "5' ESTs." As used herein, the term "purified" does not require absolute purity; rather, it is intended as a relative definition. Individual 5' EST clones isolated from a cDNA library have been conventionally purified to electrophoretic homogeneity. The sequences obtained from these clones could not be obtained directly either from the library or from total human DNA. The cDNA clones are not naturally occurring as such, but rather are obtained via manipulation of a partially purified naturally occurring substance (messenger RNA). The conversion of mRNA into a cDNA library involves the creation of a synthetic substance (cDNA) and pure individual cDNA clones can be isolated from the synthetic library by clonal selection. Thus, creating a cDNA library from messenger RNA and subsequently isolating individual clones from that library results in an approximately  $10^4$ - $10^6$  fold purification of the native message.

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Purification of starting material or natural material to at least one order of magnitude, preferably two or three orders, and more preferably four or five orders of magnitude is expressly contemplated.

As used herein, the term "isolated" requires that the material be removed from its original environment (e.g., the natural environment if it is naturally occurring). For example, a naturally-occurring polynucleotide present in a living animal is not isolated, but the same polynucleotide, separated from some or all of the coexisting materials in the natural system, is isolated.

As used herein, the term "recombinant" means that the 5' EST is adjacent to "backbone" nucleic acid to which it is not adjacent in its natural environment. Additionally, to be "enriched" the 5' ESTs will represent 5% or more of the number of nucleic acid inserts in a population of nucleic acid backbone molecules. Backbone molecules according to the present invention include nucleic acids such as expression vectors, self-replicating nucleic acids, viruses, integrating nucleic acids, and other vectors or nucleic acids used to maintain or manipulate a nucleic acid insert of interest. Preferably, the enriched 5' ESTs represent 15% or more of the number of nucleic acid inserts in the population of recombinant backbone molecules. More preferably, the enriched 5' ESTs represent 50% or more of the number of nucleic acid inserts in the population of recombinant backbone molecules. In a highly preferred embodiment, the enriched 5' ESTs represent 90% or more of the number of nucleic acid inserts in the population of recombinant backbone molecules.

"Stringent", moderate," and "low" hybridization conditions are as defined in Example 29.

Unless otherwise indicated, a "complementary" sequence is fully complementary.

Thus, 5' ESTs in cDNA libraries in which one or more 5' ESTs make up 5% or more of the number of nucleic acid inserts in the backbone molecules are "enriched recombinant 5' ESTs" as defined herein. Likewise, 5' ESTs in a population of plasmids in which one or more 5' EST of the present invention have been inserted such that they represent 5% or more of the number of inserts in the plasmid backbone are "enriched recombinant 5' ESTs" as defined herein. However, 5' ESTs in cDNA libraries in which 5' ESTs constitute less than 5% of the number of nucleic acid inserts in the population of backbone molecules, such as libraries in

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which backbone molecules having a 5' EST insert are extremely rare, are not "enriched recombinant 5' ESTs."

In particular, the present invention relates to 5' ESTs which are derived from genes encoding secreted proteins. As used herein, a "secreted" protein is one which, when expressed in a suitable host cell, is transported across or through a membrane, including transport as a result of signal peptides in its amino acid sequence. "Secreted" proteins include without limitation proteins secreted wholly (e.g. soluble proteins), or partially (e.g. receptors) from the cell in which they are expressed. "Secreted" proteins also include without limitation proteins which are transported across the membrane of the endoplasmic reticulum.

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Such 5' ESTs include nucleic acid sequences, called signal sequences, which encode signal peptides which direct the extracellular secretion of the proteins encoded by the genes from which the 5' ESTs are derived. Generally, the signal peptides are located at the amino termini of secreted proteins.

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Secreted proteins are translated by ribosomes associated with the "rough" endoplasmic reticulum. Generally, secreted proteins are co-translationally transferred to the membrane of the endoplasmic reticulum. Association of the ribosome with the endoplasmic reticulum during translation of secreted proteins is mediated by the signal peptide. The signal peptide is typically cleaved following its co-translational entry into the endoplasmic reticulum. After delivery to the endoplasmic reticulum, secreted proteins may proceed through the Golgi apparatus. In the Golgi apparatus, the proteins may undergo post-translational modification before entering secretory vesicles which transport them across the cell membrane.

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The 5' ESTs of the present invention have several important applications. For example, they may be used to obtain and express cDNA clones which include the full protein coding sequences of the corresponding gene products, including the authentic translation start sites derived from the 5' ends of the coding sequences of the mRNAs from which the 5' ESTs are derived. These cDNAs will be referred to hereinafter as "full length cDNAs." These cDNAs may also include DNA derived from mRNA sequences upstream of the translation start site. The full length cDNA sequences may be used to express the proteins corresponding to the 5' ESTs. As discussed above, secreted proteins are therapeutically important. Thus, the proteins expressed from the cDNAs may be useful in treating or

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controlling a variety of human conditions. The 5' ESTs may also be used to obtain the corresponding genomic DNA. The term "corresponding genomic DNA" refers to the genomic DNA which encodes the mRNA from which the 5' EST was derived.

Alternatively, the 5' ESTs may be used to obtain and express extended cDNAs encoding portions of the secreted protein. The portions may comprise the signal peptides of the secreted proteins or the mature proteins generated when the signal peptide is cleaved off. The portions may also comprise polypeptides having at least 10 consecutive amino acids encoded by the extended cDNAs or full length cDNAs. Alternatively, the portions may comprise at least 15 consecutive amino acids encoded by the extended cDNAs or full length cDNAs. In some embodiments, the portions may comprise at least 25 consecutive amino acids encoded by the extended cDNAs or full length cDNAs. In other embodiments, the portions may comprise at least 40 amino acids encoded by the extended cDNAs or full length cDNAs.

Antibodies which specifically recognize the entire secreted proteins encoded by the extended cDNAs, full length cDNAs, or fragments thereof having at least 10 consecutive amino acids, at least 15 consecutive amino acids, at least 25 consecutive amino acids, or at least 40 consecutive amino acids may also be obtained as described below. Antibodies which specifically recognize the mature protein generated when the signal peptide is cleaved may also be obtained as described below. Similarly, antibodies which specifically recognize the signal peptides encoded by the extended cDNAs or full length cDNAs may also be obtained.

In some embodiments, the extended cDNAs obtained using the 5' ESTs include the signal sequence. In other embodiments, the extended cDNAs obtained using the 5' ESTs may include the full coding sequence for the mature protein (*i.e.* the protein generated when the signal polypeptide is cleaved off). In addition, the extended cDNAs obtained using the 5' ESTs may include regulatory regions upstream of the translation start site or downstream of the stop codon which control the amount, location, or developmental stage of gene expression.

As discussed above, secreted proteins are therapeutically important. Thus, the proteins expressed from the extended cDNAs or full length cDNAs obtained using the 5' ESTs may be useful in treating or controlling a variety of human conditions.

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The 5' ESTs (or cDNAs or genomic DNAs obtained therefrom) may be used in forensic procedures to identify individuals or in diagnostic procedures to identify individuals having genetic diseases resulting from abnormal expression of the genes corresponding to the 5' ESTs. In addition, the present invention is useful for constructing a high resolution map of the human chromosomes.

The present invention also relates to secretion vectors capable of directing the secretion of a protein of interest. Such vectors may be used in gene therapy strategies in which it is desired to produce a gene product in one cell which is to be delivered to another location in the body. Secretion vectors may also facilitate the purification of desired proteins.

The present invention also relates to expression vectors capable of directing the expression of an inserted gene in a desired spatial or temporal manner or at a desired level. Such vectors may include sequences upstream of the 5' ESTs, such as promoters or upstream regulatory sequences.

Finally, the present invention may also be used for gene therapy to control or treat genetic diseases. Signal peptides may also be fused to heterologous proteins to direct their extracellular secretion.

Bacterial clones containing Bluescript plasmids having inserts containing the 5' ESTs of the present invention (SEQ ID NOs: 38-270 are presently stored at 80°C in 4% (v/v) glycerol in the inventor's laboratories under the designations listed next to the SEQ ID NOs in II). The inserts may be recovered from the deposited materials by growing the appropriate clones on a suitable medium. The Bluescript DNA can then be isolated using plasmid isolation procedures familiar to those skilled in the art such as alkaline lysis minipreps or large scale alkaline lysis plasmid isolation procedures. If desired the plasmid DNA may be further enriched by centrifugation on a cesium chloride gradient, size exclusion chromatography, or anion exchange chromatography. The plasmid DNA obtained using these procedures may then be manipulated using standard cloning techniques familiar to those skilled in the art. Alternatively, a PCR can be done with primers designed at both ends of the EST insertion. The PCR product which corresponds to the 5' EST can then be manipulated using standard cloning techniques familiar to those skilled in the art.

One aspect of the present invention is a purified or isolated nucleic acid having the sequence of one of SEQ ID NOs: 38-270 or having a sequence complementary thereto. In one embodiment, the nucleic acid is recombinant.

Another aspect of the present invention is a purified or isolated nucleic acid comprising at least 10 consecutive bases of the sequence of one of SEQ ID NOs: 38-270 or one of the sequences complementary thereto.

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Yet another aspect of the present invention is a purified or isolated nucleic acid comprising at least 15 consecutive bases of one of the sequences of SEQ ID NOs: 38-270 or one of the sequences complementary thereto. In one embodiment, the nucleic acid is recombinant.

A further aspect of the present invention is a purified or isolated nucleic acid of at least 15 bases capable of hybridizing under stringent conditions to the sequence of one of SEQ ID NOs: 38-270 or one of the sequences complementary to the sequences of SEQ ID NOs: 38-270. In one embodiment, the nucleic acid is recombinant.

Another aspect of the present invention is a purified or isolated nucleic acid encoding a human gene product, said human gene product having a sequence partially encoded by one of the sequences of SEQ ID NO: 38-270.

Still another aspect of the present invention is a method of making a cDNA encoding a human secretory protein, said human secretory protein being partially encoded by one of SEQ ID NOs 38-270, comprising the steps of contacting a collection of mRNA molecules from human cells with a primer comprising at least 15 consecutive nucleotides of a sequence complementary to one of SEQ ID NOs: 38-270; hybridizing said primer to an mRNA in said collection that encodes said protein; reverse transcribing said hybridized primer to make a first cDNA strand from said mRNA; making a second cDNA strand complementary to said first cDNA strand; and isolating the resulting cDNA encoding said protein comprising said first cDNA strand and said second cDNA strand.

Another aspect of the invention is an isolated or purified cDNA encoding a human secretory protein, said human secretory protein comprising the protein encoded by one of SEQ ID NOs 38-270 or a fragment thereof of at least 10 amino acids, said cDNA being obtainable by the method described in the preceding paragraph. In one embodiment, the

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cDNA comprises the full protein coding sequence of said protein which sequence is partially included in one of the sequences of SEQ ID NOs: 38-270.

Another aspect of the present invention is a method of making a cDNA encoding a human secretory protein that is partially encoded by one of SEQ ID NOs 38-270, comprising the steps of obtaining a cDNA comprising one of the sequences of SEQ ID NOs: 38-270, contacting said cDNA with a detectable probe comprising at least 15 consecutive nucleotides of said sequence of SEQ ID NO: 38-270 or a sequence complementary thereto under conditions which permit said probe to hybridize to said cDNA; identifying a cDNA which hybridizes to said detectable probe; and isolating said cDNA which hybridizes to said probe.

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Another aspect of the present invention is an isolated or purified cDNA encoding a human secretory protein, said human secretory protein comprising the protein encoded by one of SEQ ID NOs 38-270 or a fragment thereof of at least 10 amino acids, said cDNA being obtainable by the method described in the preceding paragraph. In one embodiment, the cDNA comprises the full protein coding sequence partially included in one of the sequences of SEQ ID NOs: 38-270.

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Another aspect of the present invention is a method of making a cDNA comprising one of the sequence of SEQ ID NOs: 38-270, comprising the steps of contacting a collection of mRNA molecules from human cells with a first primer capable of hybridizing to the polyA tail of said mRNA; hybridizing said first primer to said polyA tail; reverse transcribing said mRNA to make a first cDNA strand; making a second cDNA strand complementary to said first cDNA strand using at least one primer comprising at least 15 nucleotides of one of the sequences of SEQ ID NOs 38-270; and isolating the resulting cDNA comprising said first cDNA strand and said second cDNA strand.

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Another aspect of the present invention is an isolated or purified cDNA encoding a human secretory protein, said human secretory protein comprising the protein encoded by one of SEQ ID NOs 38-270 or a fragment thereof of at least 10 amino acids, said cDNA being obtainable by the method described in the preceding paragraph. In one embodiment, the cDNA comprises the full protein coding sequence partially included in one of the sequences of SEQ ID NOs: 38-270.

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In one embodiment of the method described in the two paragraphs above, the second cDNA strand is made by contacting said first cDNA strand with a first pair of primers, said

first pair of primers comprising a second primer comprising at least 15 consecutive nucleotides of one of the sequences of SEQ ID NOs 38-270 and a third primer having a sequence therein which is included within the sequence of said first primer; performing a first polymerase chain reaction with said first pair of nested primers to generate a first PCR product; contacting said first PCR product with a second pair of primers, said second pair of primers comprising a fourth primer, said fourth primer comprising at least 15 consecutive nucleotides of said sequence of one of SEQ ID NOs: 38-270, and a fifth primer, said fourth and fifth primers being capable of hybridizing to sequences within said first PCR product; and performing a second polymerase chain reaction, thereby generating a second PCR product.

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One aspect of the present invention is an isolated or purified cDNA encoding a human secretory protein, said human secretory protein comprising the protein encoded by one of SEQ ID NOs 38-270, or a fragment thereof of at least 10 amino acids, said cDNA being obtainable by the method of the preceding paragraph. In one embodiment, the cDNA comprises the full protein coding sequence partially included in one of the sequences of SEQ ID NOs: 38-270.

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Another aspect of the present invention is the method described four paragraphs above in which the second cDNA strand is made by contacting said first cDNA strand with a second primer comprising at least 15 consecutive nucleotides of the sequences of SEQ ID NOs: 38-270; hybridizing said second primer to said first strand cDNA; and extending said hybridized second primer to generate said second cDNA strand.

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Another aspect of the present invention is an isolated or purified cDNA encoding a human secretory protein, said human secretory protein comprising the protein partially encoded by one of SEQ ID NOs 38-270 or comprising a fragment thereof of at least 10 amino acids, said cDNA being obtainable by the method described in the preceding paragraph. In one embodiment, the cDNA comprises the full protein coding sequence partially included in of one of the sequences of SEQ ID NOs: 38-270.

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Another aspect of the present invention is a method of making a protein comprising one of the sequences of SEQ ID NOs: 271-503, comprising the steps of obtaining a cDNA encoding the full protein sequence partially included in one of the sequences of sequence of SEQ ID NOs: 38-270; inserting said cDNA in an expression vector such that said cDNA is

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operably linked to a promoter; introducing said expression vector into a host cell whereby said host cell produces the protein encoded by said cDNA; and isolating said protein.

Another aspect of the present invention is an isolated protein obtainable by the method described in the preceding paragraph.

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Another aspect of the present invention is a method of obtaining a promoter DNA comprising the steps of obtaining DNAs located upstream of the nucleic acids of SEQ ID NOs: 38-270 or the sequences complementary thereto; screening said upstream DNAs to identify a promoter capable of directing transcription initiation; and isolating said DNA comprising said identified promoter. In one embodiment, the obtaining step comprises chromosome walking from said nucleic acids of SEQ ID NOs: 38-270 or sequences complementary thereto. In another embodiment, the screening step comprises inserting said upstream sequences into a promoter reporter vector. In another embodiment, the screening step comprises identifying motifs in said upstream DNAs which are transcription factor binding sites or transcription start sites.

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Another aspect of the present invention is an isolated promoter obtainable by the method described above.

Another aspect of the present invention is an isolated or purified protein comprising one of the sequences of SEQ ID NOs: 271-503.

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Another aspect of the present invention is the inclusion of at least one of the sequences of SEQ ID NOs: 38-270, or one of the sequences complementary to the sequences of SEQ ID NOs: 38-270, or a fragment thereof of at least 15 consecutive nucleotides in an array of discrete ESTs or fragments thereof of at least 15 nucleotides in length. In one embodiment, the array includes at least two of the sequences of SEQ ID NOs: 38-270, the sequences complementary to the sequences of SEQ ID NOs: 38-270, or fragments thereof of at least 15 consecutive nucleotides. In another embodiment, the array includes at least five of the sequences of SEQ ID NOs: 38-270, the sequences complementary to the sequences of SEQ ID NOs: 38-270, or fragments thereof of at least 15 consecutive nucleotides.

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Another aspect of the present invention is a promoter having a sequence selected from the group consisting of SEQ ID NOs: 31, 34, and 37.

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## **Brief Description of the Drawings**

Figure 1 is a summary of a procedure for obtaining cDNAs which have been selected to include the 5' ends of the mRNAs from which they derived.

Figure 2 shows the distribution of Von Heijne scores for 5' ESTs in each of the categories described herein and the probability that these 5' ESTs encode a signal peptide.

Figure 3 summarizes a general method used to clone and sequence extended cDNAs containing sequences adjacent to 5' ESTs.

Figure 4 (description of promoters structure isolated from SignalTag 5' ESTs) provides a schematic description of promoters isolated and the way they are assembled with the corresponding 5' tags.

#### **Detailed Description of the Preferred Embodiment**

Table IV is an analysis of the 43 amino acids located at the N terminus of all human SwissProt proteins to determine the frequency of false positives and false negatives using the techniques for signal peptide identification described herein.

Table V shows the distribution of 5' ESTs in each category described herein and the number of 5' ESTs in each category having a given minimum Von Heijne's score.

Table VI shows the distribution of 5' ESTs in each category described herein with respect to the tissue from which the 5' ESTs of the corresponding mRNA were obtained.

Table VII describes the transcription factor binding sites present in each of these promoters.

#### I. General Methods for Obtaining 5' ESTs derived from mRNAs with intact 5' ends

In order to obtain the 5' ESTs of the present invention, mRNAs with intact 5' ends must be obtained. Currently, there are two approaches for obtaining such mRNAs with intact 5' ends as described below: either chemical (1) or enzymatic (2).

#### 1. Chemical Methods for Obtaining mRNAs having Intact 5' Ends

One of these approaches is a chemical modification method involving derivatization of the 5' ends of the mRNAs and selection of the derivatized mRNAs. The 5' ends of eukaryotic mRNAs possess a structure referred to as a "cap" which comprises a guanosine

methylated at the 7 position. The cap is joined to the first transcribed base of the mRNA by a 5′, 5′-triphosphate bond. In some instances, the 5′ guanosine is methylated in both the 2 and 7 positions. Rarely, the 5′ guanosine is trimethylated at the 2, 7 and 7 positions. In the chemical method for obtaining mRNAs having intact 5′ ends, the 5′ cap is specifically derivatized and coupled to a reactive group on an immobilizing substrate. This specific derivatization is based on the fact that only the ribose linked to the methylated guanosine at the 5′ end of the mRNA and the ribose linked to the base at the 3′ terminus of the mRNA, possess 2′, 3′-cis diols.

Optionally, the 2', 3'-cis diol of the 3' terminal ribose may be chemically modified, substituted, converted, or eliminated, leaving only the ribose linked to the methylated guanosine at the 5' end of the mRNA with a 2', 3'-cis diol. A variety of techniques are available for eliminating the 2', 3'-cis diol on the 3' terminal ribose. For example, controlled alkaline hydrolysis may be used to generate mRNA fragments in which the 3' terminal ribose is a 3'-phosphate, 2'-phosphate or (2', 3')-cyclophosphate. Thereafter, the fragment which includes the original 3' ribose may be eliminated from the mixture through chromatography on an oligodT column. Alternatively, a base which lacks the 2', 3'-cis diol may be added to the 3' end of the mRNA using an RNA ligase such as T4 RNA ligase. Example 1 below describes a method for ligation of a nucleoside diphosphate to the 3' end of messenger RNA.

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#### **EXAMPLE 1**

## Ligation of the Nucleoside Diphosphate pCp to the 3' End of mRNA.

One µg of RNA was incubated in a final reaction medium of 10 µl in the presence of 5 U of T<sub>4</sub> phage RNA ligase in the buffer provided by the manufacturer (Gibco - BRL), 40 U of the RNase inhibitor RNasin (Promega) and, 2 µl of <sup>32</sup>pCp (Amersham #PB 10208). The incubation was performed at 37°C for 2 hours or overnight at 7-8°C.

Following modification or elimination of the 2', 3'-cis diol at the 3' ribose, the 2', 3'-cis diol present at the 5' end of the mRNA may be oxidized using reagents such as NaBH, NaBH<sub>3</sub>CN, or sodium periodate, thereby converting the 2', 3'-cis diol to a dialdehyde. Example 2 describes the oxidation of the 2', 3'-cis diol at the 5' end of the mRNA with

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#### **EXAMPLE 2**

## Oxidation of 2', 3'-cis diol at the 5' End of the mRNA with Sodium Periodate

0.1 OD unit of either a capped oligoribonucleotide of 47 nucleotides (including the cap) or an uncapped oligoribonucleotide of 46 nucleotides were treated as follows. The oligoribonucleotides were produced by *in vitro* transcription using the transcription kit "AmpliScribe T7" (Epicentre Technologies). As indicated below, the DNA template for the RNA transcript contained a single cytosine. To synthesize the uncapped RNA, all four NTPs were included in the *in vitro* transcription reaction. To obtain the capped RNA, GTP was replaced by an analogue of the cap, m7G(5')ppp(5')G. This compound, recognized by the polymerase, was incorporated into the 5' end of the nascent transcript during the initiation of transcription but was not incorporated during the extension step. Consequently, the resulting RNA contained a cap at its 5' end. The sequences of the oligoribonucleotides produced by the *in vitro* transcription reaction were:

+Cap:

5'm7GpppGCAUCCUACUCCAUCCAAUUCCACCCUAACUCCUCCCAUCUCCAC3' (SEQ ID NO:1)

-Cap:

5'-pppGCAUCCUACUCCAUCCAAUUCCACCCUAACUCCUCCCAUCUCCAC-3' (SEQ ID NO:2)

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The oligoribonucleotides were dissolved in 9  $\mu$ l of acetate buffer (0.1 M sodium acetate, pH 5.2) and 3  $\mu$ l of freshly prepared 0.1 M sodium periodate solution. The mixture was incubated for 1 hour in the dark at 4°C or room temperature. Thereafter, the reaction was stopped by adding 4  $\mu$ l of 10% ethylene glycol. The product was ethanol precipitated, resuspended in at least 10  $\mu$ l of water or appropriate buffer and dialyzed against water.

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The resulting aldehyde groups may then be coupled to molecules having a reactive amine group, such as hydrazine, carbazide, thiocarbazide or semicarbazide groups, in order to facilitate enrichment of the 5' ends of the mRNAs. Molecules having reactive amine groups which are suitable for use in selecting mRNAs having intact 5' ends include avidin, proteins, antibodies, vitamins, ligands capable of specifically binding to receptor molecules, or oligonucleotides. Example 3 below describes the coupling of the resulting dialdehyde to biotin.

#### **EXAMPLE 3**

## Coupling of the Dialdehyde at the 5' End of Transcripts with Biotin

The oxidation product obtained in Example 2 was dissolved in 50  $\mu$ l of sodium acetate at a pH between 5 and 5.2 and 50  $\mu$ l of freshly prepared 0.02 M solution of biotin hydrazide in a methoxyethanol/water mixture (1:1) of formula:

In the compound used in these experiments, n=5. However, it will be appreciated that other commercially available hydrazides may also be used, such as molecules of the above formula in which n varies from 0 to 5. The mixture was then incubated for 2 hours at 37°C, precipitated with ethanol and dialyzed against distilled water. Example 4 demonstrates the specificity of the biotinylation reaction.

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#### **EXAMPLE 4**

### Specificity of Biotinylation of Capped Transcripts

The specificity of the biotinylation for capped mRNAs was evaluated by gel electrophoresis of the following samples:

- Sample 1. The 46 nucleotide uncapped *in vitro* transcript prepared as in Example 2 and labeled with <sup>32</sup>pCp as described in Example 1.
- Sample 2. The 46 nucleotide uncapped *in vitro* transcript prepared as in Example 2, labeled with <sup>32</sup>pCp as described in Example 1, treated with the oxidation reaction of Example 2, and subjected to the biotinylation conditions of Example 3.
- Sample 3. The 47 nucleotide capped *in vitro* transcript prepared as in Example 2 and labeled with <sup>32</sup>pCp as described in Example 1.

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Sample 4. The 47 nucleotide capped *in vitro* transcript prepared as in Example 2, labeled with <sup>32</sup>pCp as described in Example 1, treated with the oxidation reaction of Example 2, and subjected to the biotinylation conditions of Example 3.

Samples 1 and 2 had identical migration rates, demonstrating that the uncapped RNAs were not oxidized and biotinylated. Sample 3 migrated more slowly than Samples 1 and 2, while Sample 4 exhibited the slowest migration. The difference in migration of the RNAs in Samples 3 and 4 demonstrates that the capped RNAs were specifically biotinylated.

In some cases, mRNAs having intact 5' ends may be enriched by binding the molecule containing a reactive amine group to a suitable solid phase substrate such as the inside of the vessel containing the mRNAs, magnetic beads, chromatography matrices, or nylon or nitrocellulose membranes. For example, where the molecule having a reactive amine group is biotin, the solid phase substrate may be coupled to avidin or streptavidin. Alternatively, where the molecule having the reactive amine group is an antibody or receptor ligand, the solid phase substrate may be coupled to the cognate antigen or receptor. Finally, where the molecule having a reactive amine group comprises an oligonucleotide, the solid phase substrate may comprise a complementary oligonucleotide.

The mRNAs having intact 5' ends may be released from the solid phase following the enrichment procedure. For example, where the dialdehyde is coupled to biotin hydrazide and the solid phase comprises streptavidin, the mRNAs may be released from the solid phase by simply heating to 95 degrees Celsius in 2% SDS. In some methods, the molecule having a reactive amine group may also be cleaved from the mRNAs having intact 5' ends following enrichment. Example 5 describes the capture of biotinylated mRNAs with streptavidin coated beads and the release of the biotinylated mRNAs from the beads following enrichment.

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#### **EXAMPLE 5**

### Capture and Release of Biotinylated mRNAs Using Streptavidin Coated Beads

The streptavidin coated magnetic beads were prepared according to the manufacturer's instructions (CPG Inc., USA). The biotinylated mRNAs were added to a hybridization buffer (1.5 M NaCl, pH 5 - 6). After incubating for 30 minutes, the unbound and nonbiotinylated material was removed. The beads were then washed several times in

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water with 1% SDS. The beads thus obtained were incubated for 15 minutes at 95°C in water containing 2% SDS.

Example 6 demonstrates the efficiency with which biotinylated mRNAs were recovered from the streptavidin coated beads.

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#### **EXAMPLE 6**

## Efficiency of Recovery of Biotinylated mRNAs

The efficiency of the recovery procedure was evaluated as follows. Capped RNAs were labeled with <sup>32</sup>pCp, oxidized, biotinylated and bound to streptavidin coated beads as described above. Subsequently, the bound RNAs were incubated for 5, 15 or 30 minutes at 95°C in the presence of 2% SDS.

The products of the reaction were analyzed by electrophoresis on 12% polyacrylamide gels under denaturing conditions (7 M urea). The gels were subjected to autoradiography. During this manipulation, the hydrazone bonds were not reduced.

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Increasing amounts of nucleic acids were recovered as incubation times in 2% SDS increased, demonstrating that biotinylated mRNAs were efficiently recovered.

In an alternative method for obtaining mRNAs having intact 5' ends, an oligonucleotide which has been derivatized to contain a reactive amine group is specifically coupled to mRNAs having an intact cap. Preferably, the 3' end of the mRNA is blocked prior to the step in which the aldehyde groups are joined to the derivatized oligonucleotide, as described above, so as to prevent the derivatized oligonucleotide from being joined to the 3' end of the mRNA using T4 RNA ligase as described in example 1. However, as discussed above, blocking the 3' end of the mRNA is an optional step. Derivatized oligonucleotides may be prepared as described in Example 7.

#### **EXAMPLE 7**

#### Derivatization of Oligonucleotides

An oligonucleotide phosphorylated at its 3' end was converted to a 3' hydrazide in 3' by treatment with an aqueous solution of hydrazine or of dihydrazide of the formula  $H_2N(R1)NH_2$  at about 1 to 3 M, and at pH 4.5 at a temperature of 8°C overnight. This

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incubation was performed in the presence of a carbodiimide type agent soluble in water such as 1-ethyl-3-(3-dimethylaminopropyl)carbodiimide at a final concentration of 0.3 M.

The derivatized oligonucleotide was then separated from the other agents and products using a standard technique for isolating oligonucleotides.

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As discussed above, the mRNAs to be enriched may be treated to eliminate the 3' OH groups which may be present thereon. This may be accomplished by enzymatic ligation of sequences lacking a 3' OH, such as pCp, as described in Example 1. Alternatively, the 3' OH groups may be eliminated by alkaline hydrolysis as described in Example 8 below.

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#### **EXAMPLE 8**

## Elimination of 3' OH Groups of mRNA Using Alkaline Hydrolysis

In a total volume of 100 µl of 0.1 N sodium hydroxide, 1.5 µg mRNA is incubated for 40 to 60 minutes at 4°C. The solution is neutralized with acetic acid and precipitated with ethanol.

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Following the optional elimination of the 3' OH groups, the diol groups at the 5' ends of the mRNAs are oxidized as described below in Example 9.

#### **EXAMPLE 9**

#### Oxidation of Diols of mRNA

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Up to 1 OD unit of RNA was dissolved in 9  $\mu$ l of buffer (0.1 M sodium acetate, pH 6-7) or water and 3  $\mu$ l of freshly prepared 0.1 M sodium periodate solution. The reaction was incubated for 1 h in the dark at 4°C or room temperature. Following the incubation, the reaction was stopped by adding 4  $\mu$ l of 10% ethylene glycol. Thereafter the mixture was incubated at room temperature for 15 minutes. After ethanol precipitation, the product was resuspended in at least 10  $\mu$ l of water or appropriate buffer and dialyzed against water.

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Following oxidation of the diol groups at the 5' ends of the mRNAs, the derivatized oligonucleotide was joined to the resulting aldehydes as described in Example 10.

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#### **EXAMPLE 10**

## Ligature of Aldehydes of mRNA to Derivatized Oligonucleotides

The oxidized mRNA was dissolved in an acidic medium such as 50 µl of sodium acetate pH 4-6. Fifty µl of a solution of the derivatized oligonucleotide were added in order to obtain an mRNA:derivatized oligonucleotide ratio of 1:20. The mixture was reduced with a borohydride and incubated for 2 h at 37°C or overnight (14 h) at 10°C. The mixture was then ethanol precipitated, resuspended in 10 µl or more of water or appropriate buffer and dialyzed against distilled water. If desired, the resulting product may be analyzed using acrylamide gel electrophoresis, HPLC analysis, or other conventional techniques.

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Following the attachment of the derivatized oligonucleotide to the mRNAs, a reverse transcription reaction may be performed as described in Example 11 below.

#### **EXAMPLE 11**

#### Reverse Transcription of mRNAs Ligatured to Derivatized Oligonucleotides

An oligodeoxyribonucleotide was derivatized as follows. Three OD units of an oligodeoxyribonucleotide of sequence 5'ATCAAGAATTCGCACGAGACCATTA3' (SEQ ID NO:3) having 5'-OH and 3'-P ends were dissolved in 70 µl of a 1.5 M hydroxybenzotriazole solution, pH 5.3, prepared in dimethylformamide/water (75:25) containing 2 µg of 1-ethyl-3-(3-dimethylaminopropyl)carbodiimide. The mixture was incubated for 2 h 30 min at 22°C and then precipitated twice in LiClO<sub>4</sub>/acetone. The pellet was resuspended in 200 µl of 0.25 M hydrazine and incubated at 8°C from 3 to 14 h. Following the hydrazine reaction, the mixture was precipitated twice in LiClO<sub>4</sub>/acetone.

The messenger RNAs to be reverse transcribed were extracted from blocks of placenta having sides of 2 cm which had been stored at -80°C. The total RNA was extracted using conventional acidic phenol techniques. Oligo-dT chromatography was used to purify the mRNAs. The integrity of the mRNAs was checked by Northern-blotting.

The diol groups on 7 µg of the placental mRNAs were oxidized as described above in Example 9. The derivatized oligonucleotide was joined to the mRNAs as described in Example 10 above except that the precipitation step was replaced by an exclusion chromatography step to remove derivatized oligodeoxyribonucleotides which were not joined to mRNAs. Exclusion chromatography was performed as follows:

Ten ml of Ultrogel AcA34 (BioSepra#230151) gel, a mix of agarose and acrylamide, were equilibrated in 50 ml of a solution of 10 mM Tris pH 8.0, 300 mM NaCl, 1 mM EDTA, and 0.05% SDS. The mixture was allowed to sediment. The supernatant was eliminated and the gel was resuspended in 50 ml of buffer. This procedure was repeated 2 or 3 times.

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A glass bead (diameter 3 mm) was introduced into a 2 ml disposable pipette (length 25 cm). The pipette was filled with the gel suspension until the height of the gel stabilized at 1 cm from the top of the pipette. The column was then equilibrated with 20 ml of equilibration buffer (10 mM Tris HCl pH 7.4, 20 mM NaCl).

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Ten  $\mu$ I of the mRNA which had reacted with the derivatized oligonucleotide were mixed in 39  $\mu$ I of 10 mM urea and 2  $\mu$ I of blue-glycerol buffer, which had been prepared by dissolving 5 mg of bromophenol blue in 60% glycerol (v/v), and passing the mixture through a 0.45  $\mu$ m diameter filter.

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The column was then loaded with the mRNAs coupled to the oligonucleotide. As soon as the sample had penetrated, equilibration buffer was added. Hundred µl fractions were then collected. Derivatized oligonucleotide which had not been attached to mRNA appeared in fraction 16 and later fractions. Thus, fractions 3 to 15 were combined and precipitated with ethanol.

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To determine whether the derivatized oligonucleotide was actually linked to mRNA, one tenth of the combined fractions were spotted twice on a nylon membrane and hybridized to a radioactive probe using conventional techniques. The <sup>32</sup>P labeled probe used in these hybridizations was an oligodeoxyribonucleotide of sequence 5'TAATGGTCTCGTGCGAATTCTTGAT3' (SEQ ID NO:4) anticomplementary to the derivatized oligonucleotide. A signal observed after autoradiography, indicated that the derivatized oligonucleotide had been truly joined to the mRNA.

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The remaining nine tenth of the mRNAs which had reacted with the derivatized oligonucleotide was reverse transcribed as follows. A reverse transcription reaction was carried out with reverse transcriptase following the manufacturer's instructions and 50 pmol of nonamers with random sequence as primers.

To ensure that reverse transcription had been carried out through the cap structure, two types of experiments were performed.

In the first approach, after elimination of RNA of the cDNA:RNA heteroduplexes obtained from the reverse transcription reaction by an alkaline hydrolysis, a portion of the resulting single stranded cDNAs was spotted on a positively charged membrane and hybridized, using conventional methods, to a <sup>32</sup>P labeled probe having a sequence identical to that of the derivatized oligonucleotide. Control spots containing, 1 pmol, 100 finol, 50 finol, 10 finol and 1 finol of a control oligodeoxyribonucleotide of sequence identical to that of the derivatized oligonucleotide were included. The signal observed in the spots containing the cDNA indicated that approximately 15 finol of the derivatized oligonucleotide had been reverse transcribed. These results demonstrate that the reverse transcription can be performed through the cap and, in particular, that reverse transcriptase crosses the 5'-P-P-P-5' bond of the cap of eukaryotic messenger RNAs.

In the second type of experiment, the single stranded cDNAs obtained from the above first strand synthesis were used as template for PCR reactions. Two types of reactions were carried out. First, specific amplification of the mRNAs for alpha globin, dehydrogenase, pp15 and elongation factor E4 were carried out using the following pairs of oligodeoxyribonucleotide primers.

#### alpha-globin

GLO-S: 5'CCG ACA AGA CCA ACG TCA AGG CCG C3' (SEQ ID NO:5)
GLO-As: 5'TCA CCA GCA GGC AGT GGC TTA GGA G 3' (SEQ ID NO:6)

#### dehydrogenase

3 DH-S: 5'AGT GAT TCC TGC TAC TTT GGA TGG C3' (SEQ ID NO:7)
3 DH-As: 5'GCT TGG TCT TGT TCT GGA GTT TAG A3' (SEQ ID NO:8)

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pp15

PP15-S: 5'TCC AGA ATG GGA GAC AAG CCA ATT T3' (SEQ ID NO:9)
PP15-As: 5'AGG GAG GAG GAA ACA GCG TGA GTC C3' (SEQ ID NO:10)

30 Elongation factor E4

EFA1-S: 5'ATG GGA AAG GAA AAG ACT CAT ATC A3' (SEQ ID NO:11)

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## EF1A-As: 5'AGC AGC AAC AAT CAG GAC AGC ACA G3' (SEQ ID NO:12)

Second, non specific amplifications were also carried out with the antisense oligodeoxyribonucleotides of the pairs described above and with a primer derived from the sequence of the derivatized oligodeoxyribonucleotide (5'ATCAAGAATTCGCACGAGACCATTA3') (SEQ ID NO:13).

One twentieth of the following RT-PCR product samples were run on a 1.5% agarose gel and stained with ethidium bromide.

- Sample 1: The products of a PCR reaction using the globin primers of SEQ ID NOs 5 and 6 in the presence of cDNA.
  - Sample 2: The products of a PCR reaction using the globin primers of SEQ ID NOs 5 and 6 in the absence of added cDNA.
  - Sample 3: The products of a PCR reaction using the dehydrogenase primers of SEQ ID NOs 7 and 8 in the presence of cDNA.
- Sample 4: The products of a PCR reaction using the dehydrogenase primers of SEQ ID NOs 7 and 8 in the absence of added cDNA.
  - Sample 5: The products of a PCR reaction using the pp15 primers of SEQ ID NOs 9 and 10 in the presence of cDNA.
  - Sample 6: The products of a PCR reaction using the pp15 primers of SEQ ID NOs 9 and 10 in the absence of added cDNA.
    - Sample 7: The products of a PCR reaction using the EIF4 primers of SEQ ID NOs 11 and 12 in the presence of added cDNA.
    - Sample 8: The products of a PCR reaction using the EIF4 primers of SEQ ID NOs 11 and 12 in the absence of added cDNA.
  - A band of the size expected for the PCR product was observed only in samples 1, 3, 5 and 7, thus indicating the presence of the corresponding sequence in the cDNA population.

PCR reactions were also carried out with the antisense oligonucleotides of the globin and dehydrogenase primers (SEQ ID NOs 6 and 8) and an oligonucleotide whose sequence corresponds to that of the derivatized oligonucleotide. The presence of PCR products of the expected size in the samples equivalent to above samples 1 and 3 indicated that the derivatized oligonucleotide had been linked to mRNA.

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The above examples summarize the chemical procedure for enriching mRNAs for those having intact 5' ends as illustrated in Figure 1. Further detail regarding the chemical approaches for obtaining such mRNAs are disclosed in International Application No. WO96/34981, published November 7, 1996, which is incorporated herein by reference. Strategies based on the above chemical modifications to the 5' cap structure may be utilized to generate cDNAs selected to include the 5' ends of the mRNAs from which they derived. In one version of such procedures, the 5' ends of the mRNAs are modified as described Thereafter, a reverse transcription reaction is conducted to extend a primer above. complementary to the 5' end of the mRNA. Single stranded RNAs are eliminated to obtain a population of cDNA/mRNA heteroduplexes in which the mRNA includes an intact 5' end. The resulting heteroduplexes may be captured on a solid phase coated with a molecule capable of interacting with the molecule used to derivatize the 5' end of the mRNA. Thereafter, the strands of the heteroduplexes are separated to recover single stranded first cDNA strands which include the 5' end of the mRNA. Second strand cDNA synthesis may then proceed using conventional techniques. For example, the procedures disclosed in WO 96/34981 or in Carninci. et al., Genomics 37:327-336, 1996, the disclosures of which are incorporated herein by reference, may be employed to select cDNAs which include the sequence derived from the 5' end of the coding sequence of the mRNA.

Following ligation of the oligonucleotide tag to the 5' cap of the mRNA, a reverse transcription reaction is conducted to extend a primer complementary to the mRNA to the 5' end of the mRNA. Following elimination of the RNA component of the resulting heteroduplex using standard techniques, second strand cDNA synthesis is conducted with a primer complementary to the oligonucleotide tag.

#### 2. Enzymatic Methods for Obtaining mRNAs having Intact 5' Ends

Other techniques for selecting cDNAs extending to the 5' end of the mRNA from which they are derived are fully enzymatic. Some versions of these techniques are disclosed in Dumas Milne Edwards J.B. (Doctoral Thesis of Paris VI University, Le clonage des ADNc complets: difficultes et perspectives nouvelles. Apports pour l'etude de la regulation de l'expression de la tryptophane hydroxylase de rat, 20 Dec. 1993), EP0 625572 and Kato et al., Gene 150:243-250, 1994, the disclosures of which are incorporated herein by reference.

Briefly, in such approaches, isolated mRNA is treated with alkaline phosphatase to remove the phosphate groups present on the 5' ends of uncapped incomplete mRNAs. Following this procedure, the cap present on full length mRNAs is enzymatically removed with a decapping enzyme such as T4 polynucleotide kinase or tobacco acid pyrophosphatase. An oligonucleotide, which may be either a DNA oligonucleotide or a DNA-RNA hybrid oligonucleotide having RNA at its 3' end, is then ligated to the phosphate present at the 5' end of the decapped mRNA using T4 RNA ligase. The oligonucleotide may include a restriction site to facilitate cloning of the cDNAs following their synthesis. Example 12 below describes one enzymatic method based on the doctoral thesis of Dumas.

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#### **EXAMPLE 12**

## Enzymatic Approach for Obtaining 5' ESTs

Twenty micrograms of PolyA+ RNA were dephosphorylated using Calf Intestinal Phosphatase (Biolabs). After a phenol chloroform extraction, the cap structure of mRNA was hydrolysed using the Tobacco Acid Pyrophosphatase (purified as described by Shinshi et al.., Biochemistry 15: 2185-2190, 1976) and a hemi 5'DNA/RNA-3' oligonucleotide having an unphosphorylated 5' end, a stretch of adenosine ribophosphate at the 3' end, and an EcoRI site near the 5' end was ligated to the 5'P ends of mRNA using the T4 RNA ligase (Biolabs). Oligonucleotides suitable for use in this procedure are preferably 30 to 50 bases in length. Oligonucleotides having an unphosphorylated 5' end may be synthesized by adding a fluorochrome at the 5' end. The inclusion of a stretch of adenosine ribophosphates at the 3' end of the oligonucleotide increases ligation efficiency. It will be appreciated that the oligonucleotide may contain cloning sites other than EcoRI.

Following ligation of the oligonucleotide to the phosphate present at the 5' end of the decapped mRNA, first and second strand cDNA synthesis is carried out using conventional methods or those specified in EP0 625,572 and Kato et al. supra, and Dumas Milne Edwards, supra, the disclosures of which are incorporated herein by reference. The resulting cDNA may then be ligated into vectors such as those disclosed in Kato et al., supra or other nucleic acid vectors known to those skilled in the art using techniques such as those described in Sambrook et al., Molecular Cloning: A Laboratory Manual 2d Ed., Cold Spring Harbor Laboratory Press, 1989, the disclosure of which is incorporated herein by reference.

II. Obtention and Characterization of the 5' ESTs of the Present Invention

The 5' ESTs of the present invention were obtained using the aforementioned chemical and enzymatic approaches for enriching mRNAs for those having intact 5' ends as decribed below.

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## 1. Obtention of 5' ESTS Using mRNAs with Intact 5' Ends

First, mRNAs were prepared as described in Example 13 below.

#### **EXAMPLE 13**

#### Preparation of mRNA With Intact 5' Ends

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Total human RNAs or polyA+ RNAs derived from 29 different tissues were respectively purchased from LABIMO and CLONTECH and used to generate 44 cDNA libraries as follows. The purchased RNA had been isolated from cells or tissues using acid guanidium thiocyanate-phenol-chloroform extraction (Chomczyniski and Sacchi, Analytical Biochemistry 162:156-159, 1987). PolyA\* RNA was isolated from total RNA (LABIMO) by two passes of oligo dT chromatography, as described by Aviv and Leder, Proc. Natl. Acad. Sci. USA 69:1408-1412, 1972 in order to eliminate ribosomal RNA.

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The quality and the integrity of the polyA+ RNAs were checked. Northern blots hybridized with a globin probe were used to confirm that the mRNAs were not degraded. Contamination of the polyA mRNAs by ribosomal sequences was checked using Northern blots and a probe derived from the sequence of the 28S rRNA. Preparations of mRNAs with less than 5% of rRNAs were used in library construction. To avoid constructing libraries with RNAs contaminated by exogenous sequences (prokaryotic or fungal), the presence of bacterial 16S ribosomal sequences or of two highly expressed fungal mRNAs was examined using PCR.

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Following preparation of the mRNAs, the above described chemical and/or the enzymatic procedures for enriching mRNAs for thoses having intact 5' ends were employed to obtain 5' ESTs from various tissues. In both approaches, an oligonucleotide tag was attached to the 5' ends of the mRNAs. The oligonucleotide tag had an EcoRI site therein to facilitate later cloning procedures. To facilitate the processing of single stranded and double stranded cDNA obtained in the construction of the librairies, the same nucleotidic sequence was used to design the ligated oligonucleotide in both chemical and enzymatic approaches. Nevertheless, in the chemical procedure, the tag used was an oligodeoxyribonucleotide which was linked to the cap of the mRNA whereas in the enzymatic ligation, the tag was a chimeric hemi 5'DNA/RNA3' oligonucleotide which was ligated to the 5' end of decapped mRNA as described in example 12.

Following attachment of the oligonucleotide tag to the mRNA by either the chemical or enzymatic methods, the integrity of the mRNA was examined by performing a Northern blot with 200 to 500 ng of mRNA using a probe complementary to the oligonucleotide tag before performing the first strand synthesis as described in example 14.

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#### **EXAMPLE 14**

## cDNA Synthesis Using mRNA Templates Having Intact 5' Ends

For the mRNAs joined to oligonucleotide tags using both the chemical and enzymatic methods, first strand cDNA synthesis was performed using the Superscript II (Gibco BRL) or the Rnase H Minus M-MLV (Promega) reverse transcriptase with random nonamers as primers. In order to protect internal EcoRI sites in the cDNA from digestion at later steps in the procedure, methylated dCTP was used for first strand synthesis. After removal of RNA by an alkaline hydrolysis, the first strand of cDNA was precipitated using isopropanol in order to eliminate residual primers.

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For both the chemical and the enzymatic methods, the second strand of the cDNA was synthesized with a Klenow fragment using a primer corresponding to the 5' end of the ligated oligonucleotide described in Example 12. Preferably, the primer is 20-25 bases in length. Methylated dCTP was also used for second strand synthesis in order to protect internal EcoRI sites in the cDNA from digestion during the cloning process.

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Following cDNA synthesis, the cDNAs were cloned into pBlueScript as described in Example 15 below.

#### **EXAMPLE 15**

## Cloning of cDNAsderived from mRNA with intact 5' ends into BlueScript

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Following second strand synthesis, the ends of the cDNA were blunted with T4 DNA polymerase (Biolabs) and the cDNA was digested with EcoRI. Since methylated dCTP was

used during cDNA synthesis, the EcoRI site present in the tag was the only hemi-methylated site, hence the only site susceptible to EcoRI digestion. The cDNA was then size fractionated using exclusion chromatography (AcA, Biosepra) and fractions corresponding to cDNAs of more than 150 bp were pooled and ethanol precipitated. The cDNA was directionally cloned into the SmaI and EcoRI ends of the phagemid pBlueScript vector (Stratagene). The ligation mixture was electroporated into bacteria and propagated under appropriate antibiotic selection.

Clones containing the oligonucleotide tag attached were then selected as described in Example 16 below.

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#### **EXAMPLE 16**

## Selection of Clones Having the Oligonucleotide Tag Attached Thereto

The plasmid DNAs containing 5' EST libraries made as described above were purified (Qiagen). A positive selection of the tagged clones was performed as follows. Briefly, in this selection procedure, the plasmid DNA was converted to single stranded DNA using gene II endonuclease of the phage F1 in combination with an exonuclease (Chang et al., Gene 127:95-8, 1993) such as exonuclease III or T7 gene 6 exonuclease. The resulting single stranded DNA was then purified using paramagnetic beads as described by Fry et al., Biotechniques, 13: 124-131, 1992. In this procedure, the single stranded DNA was hybridized with a biotinylated oligonucleotide having a sequence corresponding to the 3' end of the oligonucleotide described in Example 13. Preferably, the primer has a length of 20-25 bases. Clones including a sequence complementary to the biotinylated oligonucleotide were captured by incubation with streptavidin coated magnetic beads followed by magnetic selection. After capture of the positive clones, the plasmid DNA was released from the magnetic beads and converted into double stranded DNA using a DNA polymerase such as the ThermoSequenase obtained from Amersham Pharmacia Biotech. Alternatively protocoles such as the one described in the Gene Trapper kit available from Gibco BRL may be used. The double stranded DNA was then electroporated into bacteria. The percentage of positive clones having the 5' tag oligonucleotide was estimated to typically rank between 90 and 98% using dot blot analysis.

Following electroporation, the libraries were ordered in 384-microtiter plates (MTP). A copy of the MTP was stored for future needs. Then the libraries were transferred into 96 MTP and sequenced as described below.

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#### **EXAMPLE 17**

### Sequencing of Inserts in Selected Clones

Plasmid inserts were first amplified by PCR on PE 9600 thermocyclers (Perkin-Elmer, Applied Biosystems Division, Foster City, CA), using standard SETA-A and SETA-B primers (Genset SA), AmpliTaqGold (Perkin-Elmer), dNTPs (Boehringer), buffer and cycling conditions as recommended by the Perkin-Elmer Corporation.

PCR products were then sequenced using automatic ABI Prism 377 sequencers (Perkin Elmer). Sequencing reactions were performed using PE 9600 thermocyclers with standard dye-primer chemistry and ThermoSequenase (Amersham Pharmacia Biotech). The primers used were either T7 or 21M13 (available from Genset SA) as appropriate. The primers were labeled with the JOE, FAM, ROX and TAMRA dyes. The dNTPs and ddNTPs used in the sequencing reactions were purchased from Boehringer. Sequencing buffer, reagent concentrations and cycling conditions were as recommended by Amersham.

Following the sequencing reaction, the samples were precipitated with ethanol, resuspended in formamide loading buffer, and loaded on a standard 4% acrylamide gel. Electrophoresis was performed for 2.5 hours at 3000V on an ABI 377 sequencer, and the sequence data were collected and analyzed using the ABI Prism DNA Sequencing Analysis Software, version 2.1.2.

# 2. Computer analysis of the Obtained 5' ESTs: Construction of NetGene and SignalTag databases

The sequence data from the 44 cDNA libraries made as described above were transferred to a proprietary database, where quality control and validation steps were performed. A proprietary base-caller, working using a Unix system, automatically flagged suspect peaks, taking into account the shape of the peaks, the inter-peak resolution, and the noise level. The proprietary base-caller also performed an automatic trimming. Any stretch of 25 or fewer bases having more than 4 suspect peaks was considered unreliable and was

discarded. • Sequences corresponding to cloning vector or ligation oligonucleotides were automatically removed from the EST sequences. However, the resulting EST sequences may contain 1 to 5 bases belonging to the above mentioned sequences at their 5' end. If needed, these can easily be removed on a case to case basis.

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Following sequencing as described above, the sequences of the 5' ESTs were entered in NetGene<sup>TM</sup>, a proprietary database called for storage and manipulation as described below. It will be appreciated by those skilled in the art that the data could be stored and manipulated on any medium which can be read and accessed by a computer. Computer readable media include magnetically, optically, or electronically readable media. For example, the computer readable media may be a hard disc, a floppy disc, a magnetic tape, CD-ROM, RAM, or ROM as well as other types of other media known to those skilled in the art.

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In addition, the sequence data may be stored and manipulated in a variety of data processor programs in a diversity of formats. For instance, the sequence data may be stored as text in a word processing file, such as Microsoft WORD or WORDPERFECT or as an ASCII file in a variety of database programs familiar to those of skill in the art, such as DB2, SYBASE, or ORACLE.

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The computer readable media on which the sequence information is stored may be in a personal computer, a network, a server or other computer systems known to those skilled in the art. The computer or other system preferably includes the storage media described above, and a processor for accessing and manipulating the sequence data. Once the sequence data has been stored, it may be manipulated and searched to locate those stored sequences which contain a desired nucleic acid sequence or which encode a protein having a particular functional domain. For example, the stored sequence information may be compared to other known sequences to identify homologies, motifs implicated in biological function, or structural motifs.

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Programs which may be used to search or compare the stored sequences include the MacPattern (EMBL), BLAST, and BLAST2 program series (NCBI), basic local alignment search tool programs for nucleotide (BLASTN) and peptide (BLASTX) comparisons (Altschul et al, J. Mol. Biol. 215: 403, 1990) and FASTA (Pearson and Lipman, Proc. Natl. Acad. Sci. USA 85: 2444, 1988). The BLAST programs then extend the alignments on the basis of defined match and mismatch criteria.

Motifs which may be detected using the above programs and those described in Example 28 include sequences encoding leucine zippers, helix-turn-helix motifs, glycosylation sites, ubiquitination sites, alpha helices, and beta sheets, signal sequences encoding signal peptides which direct the secretion of the encoded proteins, sequences implicated in transcription regulation such as homeoboxes, acidic stretches, enzymatic active sites, substrate binding sites, and enzymatic cleavage sites.

Before searching the cDNAs in the NetGene™ database for sequence motifs of interest, cDNAs derived from mRNAs which were not of interest were identified and eliminated from further consideration as described in Example 18 below.

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#### **EXAMPLE 18**

## Elimination of Undesired Sequences from Further Consideration

5' ESTs in the NetGene™ database which were derived from undesired sequences such as transfer RNAs, ribosomal RNAs, mitochondrial RNAs, prokaryotic RNAs, fungal RNAs, Alu sequences, L1 sequences, or repeat sequences were identified using the FASTA and BLASTN programs with the parameters listed in Table I.

To eliminate 5' ESTs encoding tRNAs from further consideration, the 5' EST sequences were compared to the sequences of 1190 known tRNAs obtained from EMBL release 38, of which 100 were human. The comparison was performed using FASTA on both strands of the 5' ESTs. Sequences having more than 80% homology over more than 60 nucleotides were identified as tRNA. Of the 144,341 sequences screened, 26 were identified as tRNAs and eliminated from further consideration.

To eliminate 5' ESTs encoding rRNAs from further consideration, the 5' EST sequences were compared to the sequences of 2497 known rRNAs obtained from EMBL release 38, of which 73 were human. The comparison was performed using BLASTN on both strands of the 5' ESTs with the parameter S=108. Sequences having more than 80% homology over stretches longer than 40 nucleotides were identified as rRNAs. Of the 144,341 sequences screened, 3,312 were identified as rRNAs and eliminated from further consideration.

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To eliminate 5' ESTs encoding mtRNAs from further consideration, the 5' EST sequences were compared to the sequences of the two known mitochondrial genomes for

which the entire genomic sequences are available and all sequences transcribed from these mitochondrial genomes including tRNAs, rRNAs, and mRNAs for a total of 38 sequences. The comparison was performed using BLASTN on both strands of the 5' ESTs with the parameter S=108. Sequences having more than 80% homology over stretches longer than 40 nucleotides were identified as mtRNAs. Of the 144,341 sequences screened, 6,110 were identified as mtRNAs and eliminated from further consideration.

Sequences which might have resulted from exogenous contaminants were eliminated from further consideration by comparing the 5' EST sequences to release 46 of the EMBL bacterial and fungal divisions using BLASTN with the parameter S=144. All sequences having more than 90% homology over at least 40 nucleotides were identified as exogenous contaminants. Of the 42 cDNA libraries examined, the average percentages of prokaryotic and fungal sequences contained therein were 0.2% and 0.5% respectively. Among these sequences, only one could be identified as a sequence specific to fungi. The others were either fungal or prokaryotic sequences having homologies with vertebrate sequences or including repeat sequences which had not been masked during the electronic comparison.

In addition, the 5' ESTs were compared to 6093 Alu sequences and 1115 L1 sequences to mask 5' ESTs containing such repeat sequences. 5' ESTs including THE and MER repeats, SSTR sequences or satellite, micro-satellite, or telomeric repeats were also eliminated from further consideration. On average, 11.5% of the sequences in the libraries contained repeat sequences. Of this 11.5%, 7% contained Alu repeats, 3.3% contained L1 repeats and the remaining 1.2% were derived from the other screened types of repetitive sequences. These percentages are consistent with those found in cDNA libraries prepared by other groups. For example, the cDNA libraries of Adams *et al.* contained between 0% and 7.4% Alu repeats depending on the source of the RNA which was used to prepare the cDNA library (Adams *et al.*, *Nature* 377:174, 1996).

The sequences of those 5' ESTs remaining after the elimination of undesirable sequences were compared with the sequences of known human mRNAs to determine the accuracy of the sequencing procedures described above.

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#### **EXAMPLE 19**

## Measurement of Sequencing Accuracy by Comparison to Known Sequences

To further determine the accuracy of the sequencing procedure described above, the sequences of 5' ESTs derived from known sequences were identified and compared to the original known sequences. First, a FASTA analysis with overhangs shorter than 5 bp on both ends was conducted on the 5' ESTs to identify those matching an entry in the public human mRNA database. The 6655 5' ESTs which matched a known human mRNA were then realigned with their cognate mRNA and dynamic programming was used to include substitutions, insertions, and deletions in the list of "errors" which would be recognized. Errors occurring in the last 10 bases of the 5' EST sequences were ignored to avoid the inclusion of spurious cloning sites in the analysis of sequencing accuracy.

This analysis revealed that the sequences incorporated in the NetGene™ database had an accuracy of more than 99.5%.

To determine the efficiency with which the above selection procedures select cDNAs which include the 5' ends of their corresponding mRNAs, the following analysis was performed.

#### **EXAMPLE 20**

## Determination of Efficiency of 5' EST Selection

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To determine the efficiency at which the above selection procedures isolated 5' ESTs which included sequences close to the 5' end of the mRNAs from which they derived, the sequences of the ends of the 5' ESTs derived from the elongation factor 1 subunit a and ferritin heavy chain genes were compared to the known cDNA sequences of these genes. Since the transcription start sites of both genes are well characterized, they may be used to determine the percentage of derived 5' ESTs which included the authentic transcription start sites.

For both genes, more than 95% of the obtained 5' ESTs actually included sequences close to or upstream of the 5' end of the corresponding mRNAs.

To extend the analysis of the reliability of the procedures for isolating 5' ESTs from ESTs in the NetGene™ database, a similar analysis was conducted using a database composed of human mRNA sequences extracted from GenBank database release 97 for

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comparison. The 5' ends of more than 85% of 5' ESTs derived from mRNAs included in the GeneBank database were located close to the 5' ends of the known sequence. As some of the mRNA sequences available in the GenBank database are deduced from genomic sequences, a 5' end matching with these sequences will be counted as an internal match. Thus, the method used here underestimates the yield of ESTs including the authentic 5' ends of their corresponding mRNAs.

The EST libraries made above included multiple 5' ESTs derived from the same mRNA. The sequences of such 5' ESTs were compared to one another and the longest 5' ESTs for each mRNA were identified. Overlapping cDNAs were assembled into continuous sequences (contigs). The resulting continuous sequences were then compared to public databases to gauge their similarity to known sequences, as described in Example 21 below.

## **EXAMPLE 21**

## Clustering of the 5' ESTs and Calculation of Novelty Indices for cDNA Libraries

For each sequenced EST library, the sequences were clustered by the 5' end. Each sequence in the library was compared to the others with BLASTN2 (direct strand, parameters S=107). ESTs with High Scoring Segment Pairs (HSPs) at least 25 bp long, having 95% identical bases and beginning closer than 10 bp from each EST 5' end were grouped. The longest sequence found in the cluster was used as representative of the group. A global clustering between libraries was then performed leading to the definition of super-contigs.

To assess the yield of new sequences within the EST libraries, a novelty rate (NR) was defined as: NR= 100 X (Number of new unique sequences found in the library/Total number of sequences from the library). Typically, novelty rating ranged between 10% and 41% depending on the tissue from which the EST library was obtained. For most of the libraries, the random sequencing of 5' EST libraries was pursued until the novelty rate reached 20%.

Following characterization as described above, the collection of 5' ESTs in NetGene<sup>TM</sup> was screened to identify those 5' ESTs bearing potential signal sequences as described in Example 22 below.

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#### **EXAMPLE 22**

## Identification of Potential Signal Sequences in 5' ESTs

The 5' ESTs in the NetGene<sup>TM</sup> database were screened to identify those having an uninterrupted open reading frame (ORF) longer than 45 nucleotides beginning with an ATG codon and extending to the end of the EST. Approximately half of the cDNA sequences in NetGene<sup>TM</sup> contained such an ORF. The ORFs of these 5' ESTs were then searched to identify potential signal motifs using slight modifications of the procedures disclosed in Von Heijne, *Nucleic Acids Res.* 14:4683-4690, 1986, the disclosure of which is incorporated herein by reference. Those 5' EST sequences encoding a stretch of at least 15 amino acid long with a score of at least 3.5 in the Von Heijne signal peptide identification matrix were considered to possess a signal sequence. Those 5' ESTs which matched a known human mRNA or EST sequence and had a 5' end more than 20 nucleotides downstream of the known 5' end were excluded from further analysis. The remaining cDNAs having signal sequences therein were included in a database called SignalTag<sup>TM</sup>.

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To confirm the accuracy of the above method for identifying signal sequences, the analysis of Example 23 was performed.

### **EXAMPLE 23**

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## Confirmation of Accuracy of Identification of Potential Signal Sequences in 5' ESTs

The accuracy of the above procedure for identifying signal sequences encoding signal peptides was evaluated by applying the method to the 43 amino acids located at the N terminus of all human SwissProt proteins. The computed Von Heijne score for each protein was compared with the known characterization of the protein as being a secreted protein or a non-secreted protein. In this manner, the number of non-secreted proteins having a score higher than 3.5 (false positives) and the number of secreted proteins having a score lower than 3.5 (false negatives) could be calculated.

Using the results of the above analysis, the probability that a peptide encoded by the 5' region of the mRNA is in fact a genuine signal peptide based on its Von Heijne's score was calculated based on either the assumption that 10% of human proteins are secreted or the

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assumption that 20% of human proteins are secreted. The results of this analysis are shown in Figure 2 and in Table IV.

Using the above method of identification of secretory proteins, 5' ESTs of the following polypeptides known to be secreted were obtained: human glucagon, gamma interferon induced monokine precursor, secreted cyclophilin-like protein, human pleiotropin, and human biotinidase precursor. Thus, the above method successfully identified those 5' ESTs which encode a signal peptide.

To confirm that the signal peptide encoded by the 5' ESTs actually functions as a signal peptide, the signal sequences from the 5' ESTs may be cloned into a vector designed for the identification of signal peptides. Such vectors are designed to confer the ability to grow in selective medium only to host cells containing a vector with an operably linked signal sequence. For example, to confirm that a 5' EST encodes a genuine signal peptide, the signal sequence of the 5' EST may be inserted upstream and in frame with a non-secreted form of the yeast invertase gene in signal peptide selection vectors such as those described in U.S. Patent No. 5,536,637, the disclosure of which is incorporated herein by reference. Growth of host cells containing signal sequence selection vectors with the correctly inserted 5' EST signal sequence confirms that the 5' EST encodes a genuine signal peptide.

Alternatively, the presence of a signal peptide may be confirmed by cloning the extended cDNAs obtained using the ESTs into expression vectors such as pXT1 (as described below in example 30), or by constructing promoter-signal sequence-reporter gene vectors which encode fusion proteins between the signal peptide and an assayable reporter protein. After introduction of these vectors into a suitable host cell, such as COS cells or NIH 3T3 cells, the growth medium may be harvested and analyzed for the presence of the secreted protein. The medium from these cells is compared to the medium from control cells containing vectors lacking the signal sequence or extended cDNA insert to identify vectors which encode a functional signal peptide or an authentic secreted protein.

Those 5' ESTs which encoded a signal peptide, as determined by the method of Example 22 above, were further grouped into four categories based on their homology to known sequences as described in Example 24 below.

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#### **EXAMPLE 24**

## Categorization of 5' ESTs Encoding a Signal Peptide

Those 5' ESTs having a sequence not matching any known vertebrate sequence nor any publicly available EST sequence were designated "new." Of the sequences in the SignalTag<sup>™</sup> database, 947 of the 5' ESTs having a Von Heijne's score of at least 3.5 fell into this category.

Those 5' ESTs having a sequence not matching any vertebrate sequence but matching a publicly known EST were designated "EST-ext", provided that the known EST sequence was extended by at least 40 nucleotides in the 5' direction. Of the sequences in the SignalTag<sup>TM</sup> database, 150 of the 5' ESTs having a Von Heijne's score of at least 3.5 fell into this category.

Those ESTs not matching any vertebrate sequence but matching a publicly known EST without extending the known EST by at least 40 nucleotides in the 5' direction were designated "EST." Of the sequences in the SignalTag<sup>TM</sup> database, 599 of the 5' ESTs having a Von Heijne's score of at least 3.5 fell into this category.

Those 5' ESTs matching a human mRNA sequence but extending the known sequence by at least 40 nucleotides in the 5' direction were designated "VERT-ext." Of the sequences in the SignalTag<sup>™</sup> database, 23 of the 5' ESTs having a Von Heijne's score of at least 3.5 fell into this category. Included in this category was a 5' EST which extended the known sequence of the human translocase mRNA by more than 200 bases in the 5' direction. A 5' EST which extended the sequence of a human tumor suppressor gene in the 5' direction was also identified.

Table V shows the distribution of 5' ESTs in each category and the number of 5' ESTs in each category having a given minimum von Heijne's score.

3. Evaluation of Spatial and Temporal Expression of mRNAs Corresponding to the 5'ESTs or Extended cDNAs

Each of the 5' ESTs was also categorized based on the tissue from which its corresponding mRNA was obtained, as described below in Example 25.

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### **EXAMPLE 25**

## Categorization of Expression Patterns

Table VI shows the distribution of 5' ESTs in each of the above defined category with respect to the tissue from which the 5'ESTs of the corresponding mRNA were obtained.

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Table II provides the sequence identification numbers of 5' EST sequences derived from brain, the categories in which these sequences fall, and the von Heijne's score of the signal peptides which they encode. The 5' EST sequences and the amino acid sequences they encode are provided in the appended sequence listings. Table III provides the sequence ID numbers of the 5' ESTs and the sequences of the signal peptides which they encode. The sequences of the 5' ESTs and the polypeptides they encode are provided in the sequence listing appended hereto.

The sequences of DNA SEQ ID NOs: 38-270 can readily be screened for any errors therein and any sequence ambiguities can be resolved by resequencing a fragment containing such errors or ambiguities on both strands. Such fragments may be obtained from the plasmids stored in the inventors' laboratory or can be isolated using the techniques described herein. Resolution of any such ambiguities or errors may be facilitated by using primers which hybridize to sequences located close to the ambiguous or erroneous sequences. For example, the primers may hybridize to sequences within 50-75 bases of the ambiguity or error. Upon resolution of an error or ambiguity, the corresponding corrections can be made in the protein sequences encoded by the DNA containing the error or ambiguity.

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In addition to categorizing the 5' ESTs with respect to their tissue of origin, the spatial and temporal expression patterns of the mRNAs corresponding to the 5' ESTs, as well as their expression levels, may be determined as described in Example 26 below. Characterization of the spatial and temporal expression patterns and expression levels of these mRNAs is useful for constructing expression vectors capable of producing a desired level of gene product in a desired spatial or temporal manner, as will be discussed in more detail below.

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Furthermore, 5' ESTs whose corresponding mRNAs are associated with disease states may also be identified. For example, a particular disease may result from the lack of expression, over expression, or under expression of an mRNA corresponding to a 5' EST. By comparing mRNA expression patterns and quantities in samples taken from healthy

individuals with those from individuals suffering from a particular disease, 5' ESTs responsible for the disease may be identified.

It will be appreciated that the results of the above characterization procedures for 5' ESTs also apply to extended cDNAs (obtainable as described below) which contain sequences adjacent to the 5' ESTs. It will also be appreciated that if desired, characterization may be delayed until extended cDNAs have been obtained rather than characterizing the ESTs themselves.

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## **EXAMPLE 26**

## Evaluation of Expression Levels and Patterns of mRNAs

## Corresponding to 5' ESTs or Extended cDNAs

Expression levels and patterns of mRNAs corresponding to 5' ESTs or extended cDNAs (obtainable as described below in example 27) may be analyzed by solution hybridization with long probes as described in International Patent Application No. WO 97/05277, the entire contents of which are hereby incorporated by reference. Briefly, a 5' EST, extended cDNA, or fragment thereof corresponding to the gene encoding the mRNA to be characterized is inserted at a cloning site immediately downstream of a bacteriophage (T3, T7 or SP6) RNA polymerase promoter to produce antisense RNA. Preferably, the 5' EST or extended cDNA has 100 or more nucleotides. The plasmid is linearized and transcribed in the presence of ribonucleotides comprising modified ribonucleotides (i.e. biotin-UTP and DIG-UTP). An excess of this doubly labeled RNA is hybridized in solution with mRNA isolated from cells or tissues of interest. The hybridizations are performed under standard stringent conditions (40-50°C for 16 hours in an 80% formamide, 0.4 M NaCl buffer, pH 7-8). The unhybridized probe is removed by digestion with ribonucleases specific for single-stranded RNA (i.e. RNases CL3, T1, Phy M, U2 or A). The presence of the biotin-UTP modification enables capture of the hybrid on a microtitration plate coated with streptavidin. The presence of the DIG modification enables the hybrid to be detected and quantified by ELISA using an anti-DIG antibody coupled to alkaline phosphatase.

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The 5' ESTs, extended cDNAs, or fragments thereof may also be tagged with nucleotide sequences for the serial analysis of gene expression (SAGE) as disclosed in UK

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Patent Application No. 2 305 241 A, the entire contents of which are incorporated by reference. In this method, cDNAs are prepared from a cell, tissue, organism or other source of nucleic acid for which gene expression patterns must be determined. The resulting cDNAs are separated into two pools. The cDNAs in each pool are cleaved with a first restriction endonuclease, called an anchoring enzyme, having a recognition site which is likely to be present at least once in most cDNAs. The fragments which contain the 5' or 3' most region of the cleaved cDNA are isolated by binding to a capture medium such as streptavidin coated beads. A first oligonucleotide linker having a first sequence for hybridization of an amplification primer and an internal restriction site for a so-called tagging endonuclease is ligated to the digested cDNAs in the first pool. Digestion with the second endonuclease produces short tag fragments from the cDNAs.

A second oligonucleotide having a second sequence for hybridization of an amplification primer and an internal restriction site is ligated to the digested cDNAs in the second pool. The cDNA fragments in the second pool are also digested with the tagging endonuclease to generate short tag fragments derived from the cDNAs in the second pool. The tags resulting from digestion of the first and second pools with the anchoring enzyme and the tagging endonuclease are ligated to one another to produce so-called ditags. In some embodiments, the ditags are concatamenzed to produce ligation products containing from 2 to 200 ditags. The tag sequences are then determined and compared to the sequences of the 5' ESTs or extended cDNAs to determine which 5' ESTs or extended cDNAs are expressed in the cell, tissue, organism, or other source of nucleic acids from which the tags were derived. In this way, the expression pattern of the 5' ESTs or extended cDNAs in the cell, tissue, organism, or other source of nucleic acids is obtained.

Quantitative analysis of gene expression may also be performed using arrays. As used herein, the term array means a one dimensional, two dimensional, or multidimensional arrangement of full length cDNAs (i.e. extended cDNAs which include the coding sequence for the signal peptide, the coding sequence for the mature protein, and a stop codon), extended cDNAs, 5' ESTs or fragments thereof of sufficient length to permit specific detection of gene expression. Preferably, the fragments are at least 15 nucleotides in length. More preferably, the fragments are at least 100 nucleotide long. More preferably, the

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fragments are more than 100 nucleotides in length. In some embodiments, the fragments may be more than 500 nucleotide long.

For example, quantitative analysis of gene expression may be performed with full length cDNAs as defined below, extended cDNAs, 5' ESTs, or fragments thereof in a complementary DNA microarray as described by Schena *et al.* (*Science* 270:467-470, 1995; *Proc. Natl. Acad. Sci. U.S.A.* 93:10614-10619, 1996). Full length cDNAs, extended cDNAs, 5' ESTs or fragments thereof are amplified by PCR and arrayed from 96-well microtiter plates onto silylated microscope slides using high-speed robotics. Printed arrays are incubated in a humid chamber to allow rehydration of the array elements and rinsed, once in 0.2% SDS for 1 min, twice in water for 1 min and once for 5 min in sodium borohydride solution. The arrays are submerged in water for 2 min at 95°C, transferred into 0.2% SDS for 1 min, rinsed twice with water, air dried and stored in the dark at 25°C.

Cell or tissue mRNA is isolated or commercially obtained and probes are prepared by a single round of reverse transcription. Probes are hybridized to 1 cm<sup>2</sup> microarrays under a 14 x 14 mm glass coverslip for 6-12 hours at 60°C. Arrays are washed for 5 min at 25°C in low stringency wash buffer (1 x SSC/0.2% SDS), then for 10 min at room temperature in high stringency wash buffer (0.1 x SSC/0.2% SDS). Arrays are scanned in 0.1 x SSC using a fluorescence laser scanning device fitted with a custom filter set. Accurate differential expression measurements are obtained by taking the average of the ratios of two independent hybridizations.

Quantitative analysis of the expression of genes may also be performed with full length cDNAs, extended cDNAs, 5' ESTs, or fragments thereof in complementary DNA arrays as described by Pietu et al.. (Genome Research 6:492-503, 1996). The full length cDNAs, extended cDNAs, 5' ESTs or fragments thereof are PCR amplified and spotted on membranes. Then, mRNAs originating from various tissues or cells are labeled with radioactive nucleotides. After hybridization and washing in controlled conditions, the hybridized mRNAs are detected by phospho-imaging or autoradiography. Duplicate experiments are performed and a quantitative analysis of differentially expressed mRNAs is then performed.

Alternatively, expression analysis of the 5' ESTs or extended cDNAs can be done through high density nucleotide arrays as described by Lockhart et al. (Nature Biotechnology

14: 1675-1680, 1996) and Sosnowsky et al. (Proc. Natl. Acad. Sci. 94:1119-1123, 1997). Oligonucleotides of 15-50 nucleotides corresponding to sequences of the 5' ESTs or extended cDNAs are synthesized directly on the chip (Lockhart et al., supra) or synthesized and then addressed to the chip (Sosnowsky et al., supra). Preferably, the oligonucleotides are about 20 nucleotides in length.

cDNA probes labeled with an appropriate compound, such as biotin, digoxigenin or fluorescent dye, are synthesized from the appropriate mRNA population and then randomly fragmented to an average size of 50 to 100 nucleotides. The said probes are then hybridized to the chip. After washing as described in Lockhart *et al*, *supra* and application of different electric fields (Sonowsky et *al*, *supra*.), the dyes or labeling compounds are detected and quantified. Duplicate hybridizations are performed. Comparative analysis of the intensity of the signal originating from cDNA probes on the same target oligonucleotide in different cDNA samples indicates a differential expression of the mRNA corresponding to the 5' EST or extended cDNA from which the oligonucleotide sequence has been designed.

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## III. Use of 5' ESTs to Clone Extended cDNAs and to Clone the Corresponding Genomic DNAs

Once 5' ESTs which include the 5' end of the corresponding mRNAs have been selected using the procedures described above, they can be utilized to isolate extended cDNAs which contain sequences adjacent to the 5' ESTs. The extended cDNAs may include the entire coding sequence of the protein encoded by the corresponding mRNA, including the authentic translation start site, the signal sequence, and the sequence encoding the mature protein remaining after cleavage of the signal peptide. Such extended cDNAs are referred to herein as "full length cDNAs." Alternatively, the extended cDNAs may include only the sequence encoding the mature protein remaining after cleavage of the signal peptide, or only the sequence encoding the signal peptide.

Example 27 below describes a general method for obtaining extended cDNAs using 5' ESTs. Example 28 below provides experimental results, using the method explained in example 27, describing several extended cDNAs including the entire coding sequence and authentic 5' end of the corresponding mRNA for several secreted proteins.

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The methods of Examples 27, 28, and 29 can also be used to obtain extended cDNAs which encode less than the entire coding sequence of the secreted proteins encoded by the genes corresponding to the 5' ESTs. In some embodiments, the extended cDNAs isolated using these methods encode at least 10 amino acids of one of the proteins encoded by the sequences of SEQ ID NOs: 38-270. In further embodiments, the extended cDNAs encode at least 20 amino acids of the proteins encoded by the sequences of SEQ ID NOs: 38-270. In further embodiments, the extended cDNAs encode at least 30 amino amino acids of the sequences of SEQ ID NOs: 38-270. In a preferred embodiment, the extended cDNAs encode a full length protein sequence, which includes the protein coding sequences of SEQ ID NOs: 38-270.

#### **EXAMPLE 27**

## General Method for Using 5' ESTs to Clone and Sequence cDNAs which Include the Entire Coding Region and the Authentic 5' End of the Corresponding mRNA

The following general method has been used to quickly and efficiently isolate extended cDNAs having the authentic 5' ends of their corresponding mRNAs as well as the full protein coding sequence and including sequence adjacent to the sequences of the 5' ESTs used to obtain them. This method may be applied to obtain extended cDNAs for any 5' EST in the NetGene<sup>TM</sup> database, including those 5' ESTs encoding polypeptides belonging to secreted proteins. The method is summarized in figure 3.

### 1. Obtention of Extended cDNAs

## a) First strand synthesis

The method takes advantage of the known 5' sequence of the mRNA. A reverse transcription reaction is conducted on purified mRNA with a poly 14dT primer containing a 49 nucleotide sequence at its 5' end allowing the addition of a known sequence at the end of the cDNA which corresponds to the 3' end of the mRNA. For example, the primer may have the following sequence: 5'-ATC GTT GAG ACT CGT ACC AGC AGA GTC ACG AGA GAG ACT ACA CGG TAC TGG TTT TTT TTT TTT TTT TTVN -3' (SEQ ID NO:14). Those skilled in the art will appreciate that other sequences may also be added to the poly dT sequence and used to prime the first strand synthesis. Using this primer and a reverse

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transcriptase such as the Superscript II (Gibco BRL) or Rnase H Minus M-MLV (Promega) enzyme, a reverse transcript anchored at the 3' polyA site of the RNAs is generated.

After removal of the mRNA hybridized to the first cDNA strand by alkaline hydrolysis, the products of the alkaline hydrolysis and the residual poly dT primer are eliminated with an exclusion column such as an AcA34 (Biosepra) matrix as explained in Example 11.

## b) Second strand synthesis

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A pair of nested primers on each end is designed based on the known 5' sequence from the 5' EST and the known 3' end added by the poly dT primer used in the first strand synthesis. Softwares used to design primers are either based on GC content and melting temperatures of oligonucleotides, such as OSP (Illier and Green, *PCR Meth. Appl.* 1:124-128, 1991), or based on the octamer frequency disparity method (Griffais *et al.*, *Nucleic Acids Res.* 19: 3887-3891, 1991) such as PC-Rare (http://bioinformatics.weizmann.ac.il/software/PC-Rare/doc/manuel.html).

Preferably, the nested primers at the 5' end are separated from one another by four to nine bases. The 5' primer sequences may be selected to have melting temperatures and specificities suitable for use in PCR.

Preferably, the nested primers at the 3' end are separated from one another by four to nine bases. For example, the nested 3' primers may have the following sequences: (5'- CCA GCA GAG TCA CGA GAG AGA CTA CAC GG -3'(SEQ ID NO:15), and 5'- CAC GAG AGA GAC TAC ACG GTA CTG G -3' (SEQ ID NO:16). These primers were selected because they have melting temperatures and specificities compatible with their use in PCR. However, those skilled in the art will appreciate that other sequences may also be used as primers.

The first PCR run of 25 cycles is performed using the Advantage Tth Polymerase Mix (Clontech) and the outer primer from each of the nested pairs. A second 20 cycle PCR using the same enzyme and the inner primer from each of the nested pairs is then performed on 1/2500 of the first PCR product. Thereafter, the primers and nucleotides are removed.

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## 2. Sequencing of Full Length Extended cDNAs or Fragments Thereof

Due to the lack of position constraints on the design of 5' nested primers compatible for PCR use using the OSP software, amplicons of two types are obtained. Preferably, the second 5' primer is located upstream of the translation initiation codon thus yielding a nested PCR product containing the whole coding sequence. Such a full length extended cDNA undergoes a direct cloning procedure as described in section a. However, in some cases, the second 5' primer is located downstream of the translation initiation codon, thereby yielding a PCR product containing only part of the ORF. Such incomplete PCR products are submitted to a modified procedure described in section b.

a) Nested PCR products containing complete ORFs

When the resulting nested PCR product contains the complete coding sequence, as predicted from the 5'EST sequence, it is cloned in an appropriate vector such as pED6dpc2, as described in section 3.

## b) Nested PCR products containing incomplete ORFs

When the amplicon does not contain the complete coding sequence, intermediate steps are necessary to obtain both the complete coding sequence and a PCR product containing the full coding sequence. The complete coding sequence can be assembled from several partial sequences determined directly from different PCR products as described in the following section.

Once the full coding sequence has been completely determined, new primers compatible for PCR use are designed to obtain amplicons containing the whole coding region. However, in such cases, 3' primers compatible for PCR use are located inside the 3' UTR of the corresponding mRNA, thus yielding amplicons which lack part of this region, *i.e.* the polyA tract and sometimes the polyadenylation signal, as illustrated in figure 3. Such full length extended cDNAs are then cloned into an appropriate vector as described in section 3.

### c) Sequencing extended cDNAs

Sequencing of extended cDNAs is performed using a Die Terminator approach with the AmpliTaq DNA polymerase FS kit available from Perkin Elmer.

In order to sequence PCR fragments, primer walking is performed using software such as OSP to choose primers and automated computer software such as ASMG (Sutton et

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al., Genome Science Technol. 1: 9-19, 1995) to construct contigs of walking sequences including the initial 5' tag using minimum overlaps of 32 nucleotides. Preferably, primer walking is performed until the sequences of full length cDNAs are obtained.

Completion of the sequencing of a given extended cDNA fragment is assessed as follows. Since sequences located after a polyA tract are difficult to determine precisely in the case of uncloned products, sequencing and primer walking processes for PCR products are interrupted when a polyA tract is identified in extended cDNAs obtained as described in case b. The sequence length is compared to the size of the nested PCR product obtained as described above. Due to the limited accuracy of the determination of the PCR product size by gel electrophoresis, a sequence is considered complete if the size of the obtained sequence is at least 70 % the size of the first nested PCR product. If the length of the sequence determined from the computer analysis is not at least 70% of the length of the nested PCR product, these PCR products are cloned and the sequence of the insertion is determined. When Northern blot data are available, the size of the mRNA detected for a given PCR product is used to finally assess that the sequence is complete. Sequences which do not fulfill the above criteria are discarded and will undergo a new isolation procedure.

Sequence data of all extended cDNAs are then transferred to a proprietary database, where quality controls and validation steps are carried out as described in example 15.

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### 3. Cloning of Full Length Extended cDNAs

The PCR product containing the full coding sequence is then cloned in an appropriate vector. For example, the extended cDNAs can be cloned into the expression vector pED6dpc2 (DiscoverEase, Genetics Institute, Cambridge, MA) as follows. pED6dpc2 vector DNA is prepared with blunt ends by performing an EcoRI digestion followed by a fill in reaction. The blunt ended vector is dephosphorylated. After removal of PCR primers and ethanol precipitation, the PCR product containing the full coding sequence or the extended cDNA obtained as described above is phosphorylated with a kinase subsequently removed by phenol-Sevag extraction and precipitation. The double stranded extended cDNA is then ligated to the vector and the resulting expression plasmid introduced into appropriate host cells.

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Since the PCR products obtained as described above are blunt ended molecules that can be cloned in either direction, the orientation of several clones for each PCR product is determined. Then, 4 to 10 clones are ordered in microtiter plates and subjected to a PCR reaction using a first primer located in the vector close to the cloning site and a second primer located in the portion of the extended cDNA corresponding to the 3' end of the mRNA. This second primer may be the antisense primer used in anchored PCR in the case of direct cloning (case a) or the antisense primer located inside the 3'UTR in the case of indirect cloning (case b). Clones in which the start codon of the extended cDNA is operably linked to the promoter in the vector so as to permit expression of the protein encoded by the extended cDNA are conserved and sequenced. In addition to the ends of cDNA inserts, approximately 50 bp of vector DNA on each side of the cDNA insert are also sequenced.

The cloned PCR products are then entirely sequenced according to the aforementioned procedure. In this case, contigation of long fragments is then performed on walking sequences that have already contigated for uncloned PCR products during primer walking. Sequencing of cloned amplicons is complete when the resulting contigs include the whole coding region as well as overlapping sequences with vector DNA on both ends.

### 4. Computer analysis of Full Length Extended cDNA

Sequences of all full length extended cDNAs are then submitted to further analysis as described below. Before searching the extended full length cDNAs for sequences of interest, extended cDNAs which are not of interest (vector RNAs, transfer RNAs, ribosomal RNAs, mitochondrial RNAs, prokaryotic RNAs and fungal RNAs) are discarded using methods essentially similar to those described for 5'ESTs in Example 18.

### a) Identification of structural features

Structural features, e.g. polyA tail and polyadenylation signal, of the sequences of full length extended cDNAs are subsequently determined as follows.

A polyA tail is defined as a homopolymeric stretch of at least 11 A with at most one alternative base within it. The polyA tail search is restricted to the last 100 nt of the sequence and limited to stretches of 11 consecutive A's because sequencing reactions are often not

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readable after such a polyA stretch. Stretches having more than 90% homology over 8 nucleotides are identified as polyA tails using BLAST2N.

To search for a polyadenylation signal, the polyA tail is clipped from the full-length sequence. The 50 bp preceding the polyA tail are first searched for the canonic polyadenylation AAUAAA signal and, if the canonic signal is not detected, for the alternative AUUAAA signal (Sheets et al., Nuc. Acids Res. 18: 5799-5805, 1990). If neither of these consensus polyadenylation signals is found, the canonic motif is searched again allowing one mismatch to account for possible sequencing errors. More than 85 % of identified polyadenylation signals of either type actually ends 10 to 30 bp from the polyA tail. Alternative AUUAAA signals represents approximately 15 % of the total number of identified polyadenylation signals.

## b) Identification of functional features

Functional features, e.g. ORFs and signal sequences, of the sequences of full length extended cDNAs were subsequently determined as follows.

The 3 upper strand frames of extended cDNAs are searched for ORFs defined as the maximum length fragments beginning with a translation intiation codon and ending with a stop codon. ORFs encoding at least 20 amino acids are preferred.

Each found ORF is then scanned for the presence of a signal peptide in the first 50 amino-acids or, where appropriate, within shorter regions down to 20 amino acids or less in the ORF, using the matrix method of von Heijne (*Nuc. Acids Res.* 14: 4683-4690, 1986), the disclosure of which is incorporated herein by reference as described in Example 22.

## c) Homology to either nucleotidic or proteic sequences

Categorization of full-length sequences may be achieved using procedures essentially similar to those described for 5'ESTs in Example 24.

Extended cDNAs prepared as described above may be subsequently engineered to obtain nucleic acids which include desired portions of the extended cDNA using conventional techniques such as subcloning, PCR, or *in vitro* oligonucleotide synthesis. For example, nucleic acids which include only the full coding sequences (*i.e.* the sequences encoding the signal peptide and the mature protein remaining after the signal peptide is cleaved off) may be

obtained using techniques known to those skilled in the art. Alternatively, conventional techniques may be applied to obtain nucleic acids which contain only the coding sequences for the mature protein remaining after the signal peptide is cleaved off or nucleic acids which contain only the coding sequences for the signal peptides.

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Similarly, nucleic acids containing any other desired portion of the coding sequences for the secreted protein may be obtained. For example, the nucleic acid may contain at least 10 consecutive bases of an extended cDNA such as one of the extended cDNAs described below. In another embodiment, the nucleic acid may contain at least 15 consecutive bases of an extended cDNA such as one of the extended cDNAs described below. Alternatively, the nucleic acid may contain at least 20 consecutive bases of an extended cDNA such as one of the extended cDNAs described below. In another embodiment, the nucleic acid may contain at least 25 consecutive bases of an extended cDNAs such as one of the extended cDNAs described below. In yet another embodiment, the nucleic acid may contain at least 40 consecutive bases of an extended cDNA such as one of the extended cDNAs described below.

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Once an extended cDNA has been obtained, it can be sequenced to determine the amino acid sequence it encodes. Once the encoded amino acid sequence has been determined, one can create and identify any of the many conceivable cDNAs that will encode that protein by simply using the degeneracy of the genetic code. For example, allelic variants or other homologous nucleic acids can be identified as described below. Alternatively, nucleic acids encoding the desired amino acid sequence can be synthesized *in vitro*.

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In a preferred embodiment, the coding sequence may be selected using the known codon or codon pair preferences for the host organism in which the cDNA is to be expressed.

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The extended cDNAs derived from the 5' ESTS of the present invention were obtained as described in Example 28 below.

#### **EXAMPLE 28**

## Characterization of cloned extended cDNAs obtained using 5' ESTs

The procedure described in Example 27 above was used to obtain the extended cDNAs derived from the 5' ESTs of the present invention in a variety of tissues. The following list provides a few examples of thus obtained extended cDNAs.

Using this approach, the full length cDNA of SEQ ID NO:17 (internal identification number 48-19-3-G1-FL1) was obtained. This cDNA falls into the "EST-ext" category described above and encodes the signal peptide MKKVLLLITAILAVAVG (SEQ ID NO: 18) having a von Heijne score of 8.2.

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The full length cDNA of SEQ ID NO:19 (internal identification number 58-34-2-E7-FL2) was also obtained using this procedure. This cDNA falls into the "EST-ext" category described above and encodes the signal peptide MWWFQQGLSFLPSALVIWTSA (SEQ ID NO:20) having a von Heijne score of 5.5.

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Another full length cDNA obtained using the procedure described above has the sequence of SEQ ID NO:21 (internal identification number 51-27-1-E8-FL1). This cDNA, falls into the "EST-ext" category described above and encodes the signal peptide MVLTTLPSANSANSPVNMPTTGPNSLSYASSALSPCLT (SEQ ID NO:22) having a von Heijne score of 5.9.

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The above procedure was also used to obtain a full length cDNA having the sequence of SEQ ID NO:23 (internal identification number 76-4-1-G5-FL1). This cDNA falls into the "EST-ext" category described above and encodes the signal peptide ILSTVTALTFAXA (SEQ ID NO:24) having a von Heijne score of 5.5.

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The full length cDNA of SEQ ID NO:25 (internal identification number 51-3-3-B10-FL3) was also obtained using this procedure. This cDNA falls into the "new" category described above and encodes a signal peptide LVLTLCTLPLAVA (SEQ ID NO:26) having a von Heijne score of 10.1.

The full length cDNA of SEQ ID NO:27 (internal identification number 58-35-2-F10-FL2) was also obtained using this procedure. This cDNA falls into the "new" category described above and encodes a signal peptide LWLLFFLVTAIHA (SEQ ID NO:28) having a von Heijne score of 10.7.

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Bacterial clones containing plasmids containing the full length cDNAs described above are presently stored in the inventor's laboratories under the internal identification numbers provided above. The inserts may be recovered from the stored materials by growing an aliquot of the appropriate bacterial clone in the appropriate medium. The plasmid DNA can then be isolated using plasmid isolation procedures familiar to those skilled in the art such as alkaline lysis minipreps or large scale alkaline lysis plasmid isolation procedures. If desired

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the plasmid DNA may be further enriched by centrifugation on a cesium chloride gradient, size exclusion chromatography, or anion exchange chromatography. The plasmid DNA obtained using these procedures may then be manipulated using standard cloning techniques familiar to those skilled in the art. Alternatively, a PCR can be done with primers designed at both ends of the cDNA insertion. The PCR product which corresponds to the cDNA can then be manipulated using standard cloning techniques familiar to those skilled in the art.

The polypeptides encoded by the extended cDNAs may be screened for the presence of known structural or functional motifs or for the presence of signatures, small amino acid sequences which are well conserved amongst the members of a protein family. The conserved regions have been used to derive consensus patterns or matrices included in the PROSITE data bank, in particular in the file prosite.dat (Release 13.0 of November 1995, located at <a href="http://expasy.hcuge.ch/sprot/prosite.html">http://expasy.hcuge.ch/sprot/prosite.html</a>. Prosite\_convert and prosite\_scan programs (<a href="http://ulrec3.unil.ch/ftpserveur/prosite\_scan">http://ulrec3.unil.ch/ftpserveur/prosite\_scan</a>) may be used to find signatures on the extended cDNAs.

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For each pattern obtained with the prosite\_convert program from the prosite.dat file, the accuracy of the detection on a new protein sequence may be assessed by evaluating the frequency of irrelevant hits on the population of human secreted proteins included in the data bank SWISSPROT. The ratio between the number of hits on shuffled proteins (with a window size of 20 amino acids) and the number of hits on native (unshuffled) proteins may be used as an index. Every pattern for which the ratio is greater than 20% (one hit on shuffled proteins for 5 hits on native proteins) may be skipped during the search with prosite\_scan. The program used to shuffle protein sequences (db\_shuffled) and the program used to determine the statistics for each pattern in the protein data banks (prosite\_statistics) are available on the ftp site <a href="http://ulrec3.unil.ch/ftpserveur/prosite\_scan">http://ulrec3.unil.ch/ftpserveur/prosite\_scan</a>.

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In addition to PCR based methods for obtaining extended cDNAs, traditional hybridization based methods may also be employed. These methods may also be used to obtain the genomic DNAs which encode the mRNAs from which the 5' ESTs were derived, mRNAs corresponding to the extended cDNAs, or nucleic acids which are homologous to extended cDNAs or 5' ESTs. Example 29 below provides examples of such methods.

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### **EXAMPLE 29**

## Methods for Obtaining cDNAs which include the Entire Coding Region and the Authentic 5'End of the Corresponding mRNA

A full length cDNA library can be made using the strategies described in Examples 13, 14, 15, and 16 above by replacing the random nonamer used in Example 14 with an oligo-dT primer. For instance, the oligonucleotide of SEQ ID NO:14 may be used.

Alternatively, a cDNA library or genomic DNA library may be obtained from a commercial source or made using techniques familiar to those skilled in the art. Such cDNA or genomic DNA librairies may be used to isolate extended cDNAs obtained from 5' EST or nucleic acids homologous to extended cDNAs or 5' EST as follows. The cDNA library or genomic DNA library is hybridized to a detectable probe comprising at least 10 consecutive nucleotides from the 5' EST or extended cDNA using conventional techniques. Preferably, the probe comprises at least 12, 15, or 17 consecutive nucleotides from the 5' EST or extended cDNA. More preferably, the probe comprises at least 20 to 30 consecutive nucleotides from the 5' EST or extended cDNA. In some embodiments, the probe comprises more than 30 nucleotides from the 5' EST or extended cDNA.

Techniques for identifying cDNA clones in a cDNA library which hybridize to a given probe sequence are disclosed in Sambrook et al., Molecular Cloning: A Laboratory Manual 2d Ed., Cold Spring Harbor Laboratory Press, 1989, the disclosure of which is incorporated herein by reference. The same techniques may be used to isolate genomic DNAs.

Briefly, cDNA or genomic DNA clones which hybridize to the detectable probe are identified and isolated for further manipulation as follows. A probe comprising at least 10 consecutive nucleotides from the 5' EST or extended cDNA is labeled with a detectable label such as a radioisotope or a fluorescent molecule. Preferably, the probe comprises at least 12, 15, or 17 consecutive nucleotides from the 5' EST or extended cDNA. More preferably, the probe comprises 20 to 30 consecutive nucleotides from the 5' EST or extended cDNA. In some embodiments, the probe comprises more than 30 nucleotides from the 5' EST or extended cDNA.

Techniques for labeling the probe are well known and include phosphorylation with polynucleotide kinase, nick translation, *in vitro* transcription, and non radioactive techniques. The cDNAs or genomic DNAs in the library are transferred to a nitrocellulose or nylon filter

and denatured. After blocking of non specific sites, the filter is incubated with the labeled probe for an amount of time sufficient to allow binding of the probe to cDNAs or genomic DNAs containing a sequence capable of hybridizing thereto.

By varying the stringency of the hybridization conditions used to identify extended cDNAs or genomic DNAs which hybridize to the detectable probe, extended cDNAS having different levels of homology to the probe can be identified and isolated as described below.

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# 1. Identification of Extended cDNA or Genomic cDNA Sequences Having a High Degree of Homology to the Labeled Probe

To identify extended cDNAs or genomic DNAs having a high degree of homology to the probe sequence, the melting temperature of the probe may be calculated using the following formulas:

For probes between 14 and 70 nucleotides in length the melting temperature (Tm) is calculated using the formula: Tm=81.5+16.6(log [Na+])+0.41(fraction G+C)-(600/N) where N is the length of the probe.

If the hybridization is carried out in a solution containing formamide, the melting temperature may be calculated using the equation Tm=81.5+16.6(log [Na+])+0.41(fraction G+C)-(0.63% formamide)-(600/N) where N is the length of the probe.

Prehybridization may be carried out in 6X SSC, 5X Denhardt's reagent, 0.5% SDS, 100 µg denatured fragmented salmon sperm DNA or 6X SSC, 5X Denhardt's reagent, 0.5% SDS, 100 µg denatured fragmented salmon sperm DNA, 50% formamide. The formulas for SSC and Denhardt's solutions are listed in Sambrook *et al.*, *supra*.

Hybridization is conducted by adding the detectable probe to the prehybridization solutions listed above. Where the probe comprises double stranded DNA, it is denatured before addition to the hybridization solution. The filter is contacted with the hybridization solution for a sufficient period of time to allow the probe to hybridize to extended cDNAs or genomic DNAs containing sequences complementary thereto or homologous thereto. For probes over 200 nucleotides in length, the hybridization may be carried out at 15-25°C below the Tm. For shorter probes, such as oligonucleotide probes, the hybridization may be conducted at 15-25°C below the Tm. Preferably, for hybridizations in 6X SSC, the

hybridization is conducted at approximately 68°C. Preferably, for hybridizations in 50% formamide containing solutions, the hybridization is conducted at approximately 42°C.

All of the foregoing hybridizations would be considered to be under "stringent" conditions.

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Following hybridization, the filter is washed in 2X SSC, 0.1% SDS at room temperature for 15 minutes. The filter is then washed with 0.1X SSC, 0.5% SDS at room temperature for 30 minutes to 1 hour. Thereafter, the solution is washed at the hybridization temperature in 0.1X SSC, 0.5% SDS. A final wash is conducted in 0.1X SSC at room temperature.

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Extended cDNAs, nucleic acids homologous to extended cDNAs or 5' ESTs, or genomic DNAs which have hybridized to the probe are identified by autoradiography or other conventional techniques.

# 2. Obtention of Extended cDNA or Genomic cDNA Sequences Having Lower Degrees of Homology to the Labeled Probe

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The above procedure may be modified to identify extended cDNAs, nucleic acids homologous to extended cDNAs, or genomic DNAs having decreasing levels of homology to the probe sequence. For example, to obtain extended cDNAs, nucleic acids homologous to extended cDNAs, or genomic DNAs of decreasing homology to the detectable probe, less stringent conditions may be used. For example, the hybridization temperature may be decreased in increments of 5°C from 68°C to 42°C in a hybridization buffer having a sodium concentration of approximately 1M. Following hybridization, the filter may be washed with 2X SSC, 0.5% SDS at the temperature of hybridization. These conditions are considered to be "moderate" conditions above 50°C and "low" conditions below 50°C.

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Alternatively, the hybridization may be carried out in buffers, such as 6X SSC, containing formamide at a temperature of 42°C. In this case, the concentration of formamide in the hybridization buffer may be reduced in 5% increments from 50% to 0% to identify clones having decreasing levels of homology to the probe. Following hybridization, the filter may be washed with 6X SSC, 0.5% SDS at 50°C. These conditions are considered to be "moderate" conditions above 25% formamide and "low" conditions below 25% formamide.

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Extended cDNAs, nucleic acids homologous to extended cDNAs, or genomic DNAs which have hybridized to the probe are identified by autoradiography.

## 3. Determination of the Degree of Homology Between the Obtained Extended cDNAs and the Labeled Probe

If it is desired to obtain nucleic acids homologous to extended cDNAs, such as allelic variants thereof or nucleic acids encoding proteins related to the proteins encoded by the extended cDNAs, the level of homology between the hybridized nucleic acid and the extended cDNA or 5' EST used as the probe may be further determined using BLAST2N; parameters may be adapted depending on the sequence length and degree of homology studied. To determine the level of homology between the hybridized nucleic acid and the extended cDNA or 5'EST from which the probe was derived, the nucleotide sequences of the hybridized nucleic acid and the extended cDNA or 5'EST from which the probe was derived are compared. For example, using the above methods, nucleic acids having at least 95% nucleic acid homology to the extended cDNA or 5'EST from which the probe was derived may be obtained and identified. Similarly, by using progressively less stringent hybridization conditions one can obtain and identify nucleic acids having at least 90%, at least 85%, at least 80% or at least 75% homology to the extended cDNA or 5'EST from which the probe was derived.

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To determine whether a clone encodes a protein having a given amount of homology to the protein encoded by the extended cDNA or 5' EST, the amino acid sequence encoded by the extended cDNA or 5' EST is compared to the amino acid sequence encoded by the hybridizing nucleic acid. Homology is determined to exist when an amino acid sequence in the extended cDNA or 5' EST is closely related to an amino acid sequence in the hybridizing nucleic acid. A sequence is closely related when it is identical to that of the extended cDNA or 5' EST or when it contains one or more amino acid substitutions therein in which amino acids having similar characteristics have been substituted for one another. Using the above methods and algorithms such as FASTA with parameters depending on the sequence length and degree of homology studied, one can obtain nucleic acids encoding proteins having at least 95%, at least 90%, at least 85%, at least 80% or at least 75% homology to the proteins encoded by the extended cDNA or 5'EST from which the probe was derived.

In addition to the above described methods, other protocols are available to obtain extended cDNAs using 5' ESTs as outlined in the following paragraphs.

Extended cDNAs may be prepared by obtaining mRNA from the tissue, cell, or organism of interest using mRNA preparation procedures utilizing polyA selection procedures or other techniques known to those skilled in the art. A first primer capable of hybridizing to the polyA tail of the mRNA is hybridized to the mRNA and a reverse transcription reaction is performed to generate a first cDNA strand.

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The first cDNA strand is hybridized to a second primer containing at least 10 consecutive nucleotides of the sequences of SEQ ID NOs 38-270. Preferably, the primer comprises at least 12, 15, or 17 consecutive nucleotides from the sequences of SEQ ID NOs 38-270. More preferably, the primer comprises 20 to 30 consecutive nucleotides from the sequences of SEQ ID NOs 38-270. In some embodiments, the primer comprises more than 30 nucleotides from the sequences of SEQ ID NOs 38-270. If it is desired to obtain extended cDNAs containing the full protein coding sequence, including the authentic translation initiation site, the second primer used contains sequences located upstream of the translation initiation site. The second primer is extended to generate a second cDNA strand complementary to the first cDNA strand. Alternatively, RT-PCR may be performed as described above using primers from both ends of the cDNA to be obtained.

Extended cDNAs containing 5' fragments of the mRNA may be prepared by hybridizing an mRNA comprising the sequence of the 5'EST for which an extended cDNA is desired with a primer comprising at least 10 consecutive nucleotides of the sequences complementary to the 5'EST and reverse transcribing the hybridized primer to make a first cDNA strand from the mRNAs. Preferably, the primer comprises at least 12, 15, or 17 consecutive nucleotides from the 5'EST. More preferably, the primer comprises 20 to 30 consecutive nucleotides from the 5'EST.

Thereafter, a second cDNA strand complementary to the first cDNA strand is synthesized. The second cDNA strand may be made by hybridizing a primer complementary to sequences in the first cDNA strand to the first cDNA strand and extending the primer to generate the second cDNA strand.

The double stranded extended cDNAs made using the methods described above are isolated and cloned. The extended cDNAs may be cloned into vectors such as plasmids or

viral vectors capable of replicating in an appropriate host cell. For example, the host cell may be a bacterial, mammalian, avian, or insect cell.

Techniques for isolating mRNA, reverse transcribing a primer hybridized to mRNA to generate a first cDNA strand, extending a primer to make a second cDNA strand complementary to the first cDNA strand, isolating the double stranded cDNA and cloning the double stranded cDNA are well known to those skilled in the art and are described in Current Protocols in Molecular Biology, John Wiley and Sons, Inc. 1997 and Sambrook et al., Molecular Cloning: A Laboratory Manual, Second Edition, Cold Spring Harbor Laboratory Press, 1989, the entire disclosures of which are incorporated herein by reference.

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Alternatively, procedures such as the one described in Example 29 may be used for obtaining full length cDNAs or extended cDNAs. In this approach, full length or extended cDNAs are prepared from mRNA and cloned into double stranded phagemids as follows. The cDNA library in the double stranded phagemids is then rendered single stranded by treatment with an endonuclease, such as the Gene II product of the phage F1, and an exonuclease (Chang et al., Gene 127:95-8, 1993). A biotinylated oligonucleotide comprising the sequence of a 5' EST, or a fragment containing at least 10 nucleotides thereof, is hybridized to the single stranded phagemids. Preferably, the fragment comprises at least 12, 15, or 17 consecutive nucleotides from the 5' EST. More preferably, the fragment comprises 20-30 consecutive nucleotides from the 5' EST. In some procedures, the fragment may comprise more than 30 consecutive nucleotides from the 5' EST.

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Hybrids between the biotinylated oligonucleotide and phagemids having inserts containing the 5' EST sequence are isolated by incubating the hybrids with streptavidin coated paramagnetic beads and retrieving the beads with a magnet (Fry et al., Biotechniques, 13: 124-131, 1992). Therafter, the resulting phagemids containing the 5' EST sequence are released from the beads and converted into double stranded DNA using a primer specific for the 5' EST sequence. Alternatively, protocoles such as the Gene Trapper kit (Gibco BRL) may be used. The resulting double stranded DNA is transformed into bacteria. Extended cDNAs containing the 5' EST sequence are identified by colony PCR or colony hybridization.

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Using any of the above described methods in section III, a plurality of extended cDNAs containing full length protein coding sequences or sequences encoding only the

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mature protein remaining after the signal peptide is cleaved off may be provided as cDNA libraries for subsequent evaluation of the encoded proteins or use in diagnostic assays as described below.

## IV. Expression of Proteins Encoded by Extended cDNAs Isolated Using 5' ESTs

Extended cDNAs containing the full protein coding sequences of their corresponding mRNAs or portions thereof, such as cDNAs encoding the mature protein, may be used to express the encoded secreted proteins or portions thereof as described in Example 30 below. If desired, the extended cDNAs may contain the sequences encoding the signal peptide to facilitate secretion of the expressed protein. It will be appreciated that a plurality of extended cDNAs containing the full protein coding sequences or portions thereof may be simultaneously cloned into expression vectors to create an expression library for analysis of the encoded proteins as described below.

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### **EXAMPLE 30**

## Expression of the Proteins Encoded by the Genes Corresponding to 5'ESTS or Portions Thereof

To express the proteins encoded by the genes corresponding to 5' ESTs (or portions thereof), full length cDNAs containing the entire protein coding region or extended cDNAs containing sequences adjacent to the 5' ESTs (or portions thereof) are obtained as described in Examples 27-29 and cloned into a suitable expression vector. If desired, the nucleic acids may contain the sequences encoding the signal peptide to facilitate secretion of the expressed protein. The nucleic acids inserted into the expression vectors may also contain sequences upstream of the sequences encoding the signal peptide, such as sequences which regulate expression levels or sequences which confer tissue specific expression.

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The nucleic acid encoding the protein or polypeptide to be expressed is operably linked to a promoter in an expression vector using conventional cloning technology. The expression vector may be any of the mammalian, yeast, insect or bacterial expression systems known in the art. Commercially available vectors and expression systems are available from a variety of suppliers including Genetics Institute (Cambridge, MA), Stratagene (La Jolla, California), Promega (Madison, Wisconsin), and Invitrogen (San Diego, California). If desired, to enhance expression and facilitate proper protein folding, the codon context and

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codon pairing of the sequence may be optimized for the particular expression organism in which the expression vector is introduced, as explained by Hatfield, et al., U.S. Patent No. 5,082,767, incorporated herein by this reference.

The cDNA cloned into the expression vector may encode the entire protein (i.e. the signal peptide and the mature protein), the mature protein (i.e. the protein created by cleaving the signal peptide off), only the signal peptide or any other portion thereof.

The following is provided as one exemplary method to express the proteins encoded by the extended cDNAs corresponding to the 5' ESTs or the nucleic acids described above. First, the methionine initiation codon for the gene and the polyA signal of the gene are identified. If the nucleic acid encoding the polypeptide to be expressed lacks a methionine to serve as the initiation site, an initiating methionine can be introduced next to the first codon of the nucleic acid using conventional techniques. Similarly, if the extended cDNA lacks a polyA signal, this sequence can be added to the construct by, for example, splicing out the polyA signal from pSG5 (Stratagene) using BgIII and SalI restriction endonuclease enzymes and incorporating it into the mammalian expression vector pXT1 (Stratagene). pXT1 contains the LTRs and a portion of the gag gene from Moloney Murine Leukemia Virus. The position of the LTRs in the construct allow efficient stable transfection. The vector includes the Herpes Simplex thymidine kinase promoter and the selectable neomycin gene. The extended cDNA or portion thereof encoding the polypeptide to be expressed is obtained by PCR from the bacterial vector using oligonucleotide primers complementary to the extended cDNA or portion thereof and containing restriction endonuclease sequences for Pst I incorporated into the 5'primer and BglII at the 5' end of the corresponding cDNA 3' primer, taking care to ensure that the extended cDNA is positioned with the poly A signal. The purified fragment obtained from the resulting PCR reaction is digested with PstI, blunt ended with an exonuclease, digested with Bgl II, purified and ligated to pXT1 containing a poly A signal and prepared for this ligation (blunt/BgIII).

The ligated product is transfected into mouse NIH 3T3 cells using Lipofectin (Life Technologies, Inc., Grand Island, New York) under conditions outlined in the product specification. Positive transfectants are selected after growing the transfected cells in 600 µg/ml G418 (Sigma, St. Louis, Missouri). Preferably the expressed protein is released into the culture medium, thereby facilitating purification.

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Alternatively, the extended cDNAs may be cloned into pED6dpc2 as described above. The resulting pED6dpc2 constructs may be transfected into a suitable host cell, such as COS 1 cells. Methotrexate resistant cells are selected and expanded. Preferably, the protein expressed from the extended cDNA is released into the culture medium thereby facilitating purification.

Proteins in the culture medium are separated by gel electrophoresis. If desired, the proteins may be ammonium sulfate precipitated or separated based on size or charge prior to electrophoresis.

As a control, the expression vector lacking a cDNA insert is introduced into host cells or organisms and the proteins in the medium are harvested. The secreted proteins present in the medium are detected using techniques familiar to those skilled in the art such as Coomassie blue or silver staining or using antibodies against the protein encoded by the extended cDNA

Antibodies capable of specifically recognizing the protein of interest may be generated using synthetic 15-mer peptides having a sequence encoded by the appropriate 5' EST, extended cDNA, or portion thereof. The synthetic peptides are injected into mice to generate antibody to the polypeptide encoded by the 5' EST, extended cDNA, or portion thereof.

Secreted proteins from the host cells or organisms containing an expression vector which contains the extended cDNA derived from a 5' EST or a portion thereof are compared to those from the control cells or organism. The presence of a band in the medium from the cells containing the expression vector which is absent in the medium from the control cells indicates that the extended cDNA encodes a secreted protein. Generally, the band corresponding to the protein encoded by the extended cDNA will have a mobility near that expected based on the number of amino acids in the open reading frame of the extended cDNA. However, the band may have a mobility different than that expected as a result of modifications such as glycosylation, ubiquitination, or enzymatic cleavage.

Alternatively, if the protein expressed from the above expression vectors does not contain sequences directing its secretion, the proteins expressed from host cells containing an expression vector with an insert encoding a secreted protein or portion thereof can be compared to the proteins expressed in control host cells containing the expression vector without an insert. The presence of a band in samples from cells containing the expression

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vector with an insert which is absent in samples from cells containing the expression vector without an insert indicates that the desired protein or portion thereof is being expressed. Generally, the band will have the mobility expected for the secreted protein or portion thereof. However, the band may have a mobility different than that expected as a result of modifications such as glycosylation, ubiquitination, or enzymatic cleavage.

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The protein encoded by the extended cDNA may be purified using standard immunochromatography techniques. In such procedures, a solution containing the secreted protein, such as the culture medium or a cell extract, is applied to a column having antibodies against the secreted protein attached to the chromatography matrix. The secreted protein is allowed to bind the immunochromatography column. Thereafter, the column is washed to remove non-specifically bound proteins. The specifically bound secreted protein is then released from the column and recovered using standard techniques.

If antibody production is not possible, the extended cDNA sequence or portion thereof may be incorporated into expression vectors designed for use in purification schemes employing chimeric polypeptides. In such strategies, the coding sequence of the extended cDNA or portion thereof is inserted in frame with the gene encoding the other half of the chimera. The other half of the chimera may be  $\beta$ -globin or a nickel binding polypeptide. A chromatography matrix having antibody to  $\beta$ -globin or nickel attached thereto is then used to purify the chimeric protein. Protease cleavage sites may be engineered between the  $\beta$ -globin gene or the nickel binding polypeptide and the extended cDNA or portion thereof. Thus, the two polypeptides of the chimera may be separated from one another by protease digestion.

One useful expression vector for generating β-globin chimerics is pSG5 (Stratagene), which encodes rabbit β-globin. Intron II of the rabbit β-globin gene facilitates splicing of the expressed transcript, and the polyadenylation signal incorporated into the construct increases the level of expression. These techniques as described are well known to those skilled in the art of molecular biology. Standard methods are published in methods texts such as Davis et al., (Basic Methods in Molecular Biology, Davis, Dibner, and Battey, ed., Elsevier Press, NY, 1986) and many of the methods are available from Stratagene, Life Technologies, Inc., or Promega. Polypeptide may additionally be produced from the construct using *in vitro* translation systems such as the *In vitro* Express<sup>TM</sup> Translation Kit (Stratagene).

Following expression and purification of the secreted proteins encoded by the 5' ESTs, extended cDNAs, or fragments thereof, the purified proteins may be tested for the ability to bind to the surface of various cell types as described in Example 31 below. It will be appreciated that a plurality of proteins expressed from these cDNAs may be included in a panel of proteins to be simultaneously evaluated for the activities specifically described below, as well as other biological roles for which assays for determining activity are available.

### **EXAMPLE 31**

## Analysis of Secreted Proteins to Determine Whether they Bind to the Cell Surface

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The proteins encoded by the 5' ESTs, extended cDNAs, or fragments thereof are cloned into expression vectors such as those described in Example 30. The proteins are purified by size, charge, immunochromatography or other techniques familiar to those skilled in the art. Following purification, the proteins are labeled using techniques known to those skilled in the art. The labeled proteins are incubated with cells or cell lines derived from a variety of organs or tissues to allow the proteins to bind to any receptor present on the cell surface. Following the incubation, the cells are washed to remove non-specifically bound protein. The labeled proteins are detected by autoradiography. Alternatively, unlabeled proteins may be incubated with the cells and detected with antibodies having a detectable label, such as a fluorescent molecule, attached thereto.

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Specificity of cell surface binding may be analyzed by conducting a competition analysis in which various amounts of unlabeled protein are incubated along with the labeled protein. The amount of labeled protein bound to the cell surface decreases as the amount of competitive unlabeled protein increases. As a control, various amounts of an unlabeled protein unrelated to the labeled protein is included in some binding reactions. The amount of labeled protein bound to the cell surface does not decrease in binding reactions containing increasing amounts of unrelated unlabeled protein, indicating that the protein encoded by the cDNA binds specifically to the cell surface.

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As discussed above, secreted proteins have been shown to have a number of important physiological effects and, consequently, represent a valuable therapeutic resource. The secreted proteins encoded by the extended cDNAs or portions thereof made according to

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Examples 27-29 may be evaluated to determine their physiological activities as described below.

### **EXAMPLE 32**

## 5 <u>Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Cytokine,</u> <u>Cell Proliferation or Cell Differentiation Activity</u>

As discussed above, secreted proteins may act as cytokines or may affect cellular proliferation or differentiation. Many protein factors discovered to date, including all known cytokines, have exhibited activity in one or more factor dependent cell proliferation assays, and hence the assays serve as a convenient confirmation of cytokine activity. The activity of a protein encoded by the extended cDNAs is evidenced by any one of a number of routine factor dependent cell proliferation assays for cell lines including, without limitation, 32D, DA2, DA1G, T10, B9, B9/11, BaF3, MC9/G, M<sup>+</sup> (preB M<sup>+</sup>), 2E8, RB5, DA1, 123, T1165, HT2, CTLL2, TF-1, Mo7c and CMK. The proteins encoded by the above extended cDNAs or portions thereof may be evaluated for their ability to regulate T cell or thymocyte proliferation in assays such as those described above or in the following references, which are incorporated herein by reference: Current Protocols in Immunology, Ed. by Coligan et al., Greene Publishing Associates and Wiley-Interscience; Takai et al. J. Immunol. 137:3494-3500, 1986., Bertagnolli et al., J. Immunol. 145:1706-1712, 1990., Bertagnolli et al., Cell. Immunol. 133:327-341, 1991; Bertagnolli, et al., J. Immunol. 149:3778-3783, 1992; Bowman et al., J. Immunol. 152:1756-1761, 1994.

In addition, numerous assays for cytokine production and/or the proliferation of spleen cells, lymph node cells and thymocytes are known. These include the techniques disclosed in *Current Protocols in Immunology*, supra 1:3.12.1-3.12.14; and Schreiber In *Current Protocols in Immunology*, supra 1:6.8.1-6.8.8.

The proteins encoded by the cDNAs may also be assayed for the ability to regulate the proliferation and differentiation of hematopoietic or lymphopoietic cells. Many assays for such activity are familiar to those skilled in the art, including the assays in the following references, which are incorporated herein by reference: Bottomly et al., In Current Protocols in Immunology., supra. 1:6.3.1-6.3.12,; deVries et al., J. Exp. Med. 173:1205-1211, 1991; Moreau et al., Nature 36:690-692, 1988; Greenberger et al., Proc. Natl. Acad. Sci. U.S.A.

80:2931-2938, 1983; Nordan, R., In Current Protocols in Immunology., supra. 1:6.6.1-6.6.5; Smith et al., Proc. Natl. Acad. Sci. U.S.A. 83:1857-1861, 1986; Bennett et al., in Current Protocols in Immunology supra 1:6.15.1; Ciarletta et al., In Current Protocols in Immunology. supra 1:6.13.1.

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The proteins encoded by the cDNAs may also be assayed for their ability to regulate T-cell responses to antigens. Many assays for such activity are familiar to those skilled in the art, including the assays described in the following references, which are incorporated herein by reference: Chapter 3 (In Vitro Assays for Mouse Lymphocyte Function), Chapter 6 (Cytokines and Their Cellular Receptors) and Chapter 7, (Immunologic Studies in Humans) in Current Protocols in Immunology supra; Weinberger et al., Proc. Natl. Acad. Sci. USA 77:6091-6095, 1980; Weinberger et al., Eur. J. Immun. 11:405-411, 1981; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988.

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Those proteins which exhibit cytokine, cell proliferation, or cell differentiation activity may then be formulated as pharmaceuticals and used to treat clinical conditions in which induction of cell proliferation or differentiation is beneficial. Alternatively, as described in more detail below, genes encoding these proteins or nucleic acids regulating the expression of these proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

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## **EXAMPLE 33**

## Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Activity as Immune System Regulators

The proteins encoded by the cDNAs may also be evaluated for their effects as immune regulators. For example, the proteins may be evaluated for their activity to influence thymocyte or splenocyte cytotoxicity. Numerous assays for such activity are familiar to those skilled in the art including the assays described in the following references, which are incorporated herein by reference: Chapter 3 (In Vitro Assays for Mouse Lymphocyte Function 3.1-3.19) and Chapter 7 (Immunologic studies in Humans) in Current Protocols in Immunology, Coligan et al., Eds, Greene Publishing Associates and Wiley-Interscience,

Herrmann et al., Proc. Natl. Acad. Sci. USA 78:2488-2492, 1981; Herrmann et al., J.

Immunol. 128:1968-1974, 1982; Handa et al., J. Immunol. 135:1564-1572, 1985; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988; Bowman et al., J. Virology 61:1992-1998; Bertagnolli et al., Cell. Immunol. 133:327-341, 1991; Brown et al., J. Immunol. 153:3079-3092, 1994.

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The proteins encoded by the cDNAs may also be evaluated for their effects on T-cell dependent immunoglobulin responses and isotype switching. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Maliszewski, *J. Immunol.* 144:3028-3033, 1990; Mond *et al.* in *Current Protocols in Immunology*, 1:3.8.1-3.8.16, *supra*.

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The proteins encoded by the cDNAs may also be evaluated for their effect on immune effector cells, including their effect on Th1 cells and cytotoxic lymphocytes. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Chapter 3 (*In Vitro* Assays for Mouse Lymphocyte Function 3.1-3.19) and Chapter 7 (Immunologic Studies in Humans) in *Current Protocols in Immunology, supra*; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988; Bertagnolli et al., J. Immunol. 149:3778-3783, 1992.

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The proteins encoded by the cDNAs may also be evaluated for their effect on dendritic cell mediated activation of naive T-cells. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Guery et al., J. Immunol. 134:536-544, 1995; Inaba et al., J. Exp. Med. 173:549-559, 1991; Macatonia et al., J. Immunol. 154:5071-5079, 1995; Porgador et al.J. Exp. Med 182:255-260, 1995; Nair et al., J. Virol. 67:4062-4069, 1993; Huang et al., Science 264:961-965, 1994; Macatonia et al.J. Exp. Med 169:1255-1264, 1989; Bhardwaj et al., Journal of Clinical Investigation 94:797-807, 1994; and Inaba et al., J. Exp. Med 172:631-640, 1990.

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The proteins encoded by the cDNAs may also be evaluated for their influence on the lifetime of lymphocytes. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Darzynkiewicz et al., Cytometry 13:795-808, 1992; Gorczyca et al., Leukemia 7:659-670, 1993; Gorczyca et al., Cancer Res. 53:1945-1951, 1993; Itoh et al., Cell 66:233-

243, 1991; Zacharchuk, J. Immunol. 145:4037-4045, 1990; Zamai et al., Cytometry 14:891-897, 1993; Gorczyca et al., Int. J. Oncol. 1:639-648, 1992.

The proteins encoded by the cDNAs may also be evaluated for their influence on early steps of T-cell commitment and development. Numerous assays for such activity are familiar to those skilled in the art, including without limitation the assays disclosed in the following references, which are incorporated herein by references: Antica et al., Blood 84:111-117, 1994; Fine et al., Cell. Immunol. 155:111-122, 1994; Galy et al., Blood 85:2770-2778, 1995; Toki et al., Proc. Nat. Acad Sci. USA 88:7548-7551, 1991.

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Those proteins which exhibit activity as immune system regulators activity may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of immune activity is beneficial. For example, the protein may be useful in the treatment of various immune deficiencies and disorders (including severe combined immunodeficiency), e.g., in regulating (up or down) growth and proliferation of T and/or B lymphocytes, as well as effecting the cytolytic activity of NK cells and other cell populations. These immune deficiencies may be genetic or be caused by viral (e.g., HIV) as well as bacterial or fungal infections, or may result from autoimmune disorders. More specifically, infectious diseases caused by viral, bacterial, fungal or other infection may be treatable using a protein encoded by extended cDNAs derived from the 5' ESTs of the present invention, including infections by HIV, hepatitis viruses, herpesviruses, mycobacteria, Leishmania spp., plamodium and various fungal infections such as candidiasis. Of course, in this regard, a protein encoded by extended cDNAs derived from the 5' ESTs of the present invention may also be useful where a boost to the immune system generally may be desirable, i.e., in the treatment of cancer.

Alternatively, proteins encoded by extended cDNAs derived from the 5' ESTs of the present invention may be used in treatment of autoimmune disorders including, for example, connective tissue disease, multiple sclerosis, systemic lupus erythematosus, rheumatoid arthritis, autoimmune pulmonary inflammation, Guillain-Barre syndrome, autoimmune thyroiditis, insulin dependent diabetes mellitis, myasthenia gravis, graft-versus-host disease and autoimmune inflammatory eye disease. Such a protein encoded by extended cDNAs derived from the 5' ESTs of the present invention may also to be useful in the treatment of allergic reactions and conditions, such as asthma (particularly allergic asthma) or other respiratory problems. Other conditions, in which immune suppression is desired (including

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for example, organ transplantation), may also be treatable using a protein encoded by extended cDNAs derived from the 5' ESTs of the present invention.

Using the proteins of the invention it may also be possible to regulate immune responses either up or down.

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Down regulation may involve inhibiting or blocking an immune response already in progress or may involve preventing the induction of an immune response. The functions of activated T-cells may be inhibited by suppressing T cell responses or by inducing specific tolerance in T cells, or both. Immunosuppression of T cell responses is generally an active non-antigen-specific process which requires continuous exposure of the T cells to the suppressive agent. Tolerance, which involves inducing non-responsiveness or anergy in T cells, is distinguishable from immunosuppression in that it is generally antigen-specific and persists after the end of exposure to the tolerizing agent. Operationally, tolerance can be demonstrated by the lack of a T cell response upon reexposure to specific antigen in the absence of the tolerizing agent.

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Down regulating or preventing one or more antigen functions (including without limitation B lymphocyte antigen functions, such as, for example, B7 costimulation), e.g., preventing high level lymphokine synthesis by activated T cells, will be useful in situations of tissue, skin and organ transplantation and in graft-versus-host disease (GVHD). For example, blockage of T cell function should result in reduced tissue destruction in tissue transplantation. Typically, in tissue transplants, rejection of the transplant is initiated through its recognition as foreign by T cells, followed by an immune reaction that destroys the transplant. The administration of a molecule which inhibits or blocks interaction of a B7 lymphocyte antigen with its natural ligand(s) on immune cells (such as a soluble, monomeric form of a peptide having B7-2 activity alone or in conjunction with a monomeric form of a peptide having an activity of another B lymphocyte antigen (e.g., B7-1, B7-3) or blocking antibody), prior to transplantation, can lead to the binding of the molecule to the natural ligand(s) on the immune cells without transmitting the corresponding costimulatory signal. Blocking B lymphocyte antigen function in this matter prevents cytokine synthesis by immune cells, such as T cells, and thus acts as an immunosuppressant. Moreover, the lack of costimulation may also be sufficient to anergize the T cells, thereby inducing tolerance in a subject. Induction of long-term tolerance by B lymphocyte antigen-blocking reagents may

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avoid the necessity of repeated administration of these blocking reagents. To achieve sufficient immunosuppression or tolerance in a subject, it may also be necessary to block the function of a combination of B lymphocyte antigens.

The efficacy of particular blocking reagents in preventing organ transplant rejection or GVHD can be assessed using animal models that are predictive of efficacy in humans. Examples of appropriate systems which can be used include allogeneic cardiac grafts in rats and xenogeneic pancreatic islet cell grafts in mice, both of which have been used to examine the immunosuppressive effects of CTLA4Ig fusion proteins *in vivo* as described in Lenschow *et al.*, *Science* 257:789-792, 1992 and Turka *et al.*, *Proc. Natl. Acad. Sci USA*, 89:11102-11105, 1992. In addition, murine models of GVHD (see Paul ed., *Fundamental Immunology*, Raven Press, New York, 1989, pp. 846-847) can be used to determine the effect of blocking B lymphocyte antigen function *in vivo* on the development of that disease.

Blocking antigen function may also be therapeutically useful for treating autoimmune diseases. Many autoimmune disorders are the result of inappropriate activation of T cells that are reactive against self tissue and which promote the production of cytokines and autoantibodies involved in the pathology of the diseases. Preventing the activation of autoreactive T cells may reduce or eliminate disease symptoms. Administration of reagents which block costimulation of T cells by disrupting receptor/ligand interactions of B lymphocyte antigens can be used to inhibit T cell activation and prevent production of autoantibodies or T cell-derived cytokines which potentially involved in the disease process. Additionally, blocking reagents may induce antigen-specific tolerance of autoreactive T cells which could lead to long-term relief from the disease. The efficacy of blocking reagents in preventing or alleviating autoimmune disorders can be determined using a number of well-characterized animal models of human autoimmune diseases. Examples include murine experimental autoimmune encephalitis, systemic lupus erythmatosis in MRL/pr/pr mice or NZB hybrid mice, murine autoimmuno collagen arthritis, diabetes mellitus in OD mice and BB rats, and murine experimental myasthenia gravis (see Paul ed., supra, pp. 840-856).

Upregulation of an antigen function (preferably a B lymphocyte antigen function), as a means of up regulating immune responses, may also be useful in therapy. Upregulation of immune responses may involve either enhancing an existing immune response or eliciting an initial immune response as shown by the following examples. For instance, enhancing an

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immune response through stimulating B lymphocyte antigen function may be useful in cases of viral infection. In addition, systemic viral diseases such as influenza, the common cold, and encephalitis might be alleviated by the administration of stimulatory form of B lymphocyte antigens systemically.

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Alternatively, antiviral immune responses may be enhanced in an infected patient by removing T cells from the patient, costimulating the T cells *in vitro* with viral antigen-pulsed APCs either expressing a peptide encoded by extended cDNAs derived from the 5' ESTs of the present invention or together with a stimulatory form of a soluble peptide encoded by extended cDNAs derived from the 5' ESTs of the present invention and reintroducing the *in vitro* primed T cells into the patient. The infected cells would now be capable of delivering a costimulatory signal to T cells *in vivo*, thereby activating the T cells.

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In another application, upregulation or enhancement of antigen function (preferably B lymphocyte antigen function) may be useful in the induction of tumor immunity. Tumor cells (e.g., sarcoma, melanoma, lymphoma, leukemia, neuroblastoma, carcinoma) transfected with a nucleic acid encoding at least one peptide encoded by extended cDNAs derived from the 5' ESTs of the present invention can be administered to a subject to overcome tumor-specific tolerance in the subject. If desired, the tumor cell can be transfected to express a combination of peptides. For example, tumor cells obtained from a patient can be transfected *ex vivo* with an expression vector directing the expression of a peptide having B7-2-like activity alone, or in conjunction with a peptide having B7-1-like activity and/or B7-3-like activity. The transfected tumor cells are returned to the patient to result in expression of the peptides on the surface of the transfected cell. Alternatively, gene therapy techniques can be used to target a tumor cell for transfection *in vivo*.

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The presence of the peptide encoded by extended cDNAs derived from the 5' ESTs of the present invention having the activity of a B lymphocyte antigen(s) on the surface of the tumor cell provides the necessary costimulation signal to T cells to induce a T cell mediated immune response against the transfected tumor cells. In addition, tumor cells which lack or which fail to reexpress sufficient amounts of MHC class I or MHC class II molecules can be transfected with nucleic acids encoding all or a portion of (e.g., a cytoplasmic-domain truncated portion) of an MHC class I  $\alpha$  chain and  $\beta_2$  microglobulin or an MHC class II  $\alpha$  chain and an MHC class II  $\beta$  chain to thereby express MHC class I or MHC class II proteins

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on the cell surface, respectively. Expression of the appropriate MHC class I or class II molecules in conjunction with a peptide having the activity of a B lymphocyte antigen (e.g., B7-1, B7-2, B7-3) induces a T cell mediated immune response against the transfected tumor cell. Optionally, a gene encoding an antisense construct which blocks expression of an MHC class II associated protein, such as the invariant chain, can also be cotransfected with a DNA encoding a peptide having the activity of a B lymphocyte antigen to promote presentation of tumor associated antigens and induce tumor specific immunity. Thus, the induction of a T cell mediated immune response in a human subject may be sufficient to overcome tumor-specific tolerance in the subject. Alternatively, as described in more detail below, genes encoding these immune system regulator proteins or nucleic acids regulating the expression of such proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

### **EXAMPLE 34**

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## Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Hematopoiesis Regulating Activity

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for their hematopoiesis regulating activity. For example, the effect of the proteins on embryonic stem cell differentiation may be evaluated. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Johansson et al. Cell. Biol. 15:141-151, 1995; Keller et al., Mol. Cell. Biol. 13:473-486, 1993; McClanahan et al., Blood 81:2903-2915, 1993.

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for their influence on the lifetime of stem cells and stem cell differentiation. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Freshney, Methylcellulose Colony Forming Assays, in Culture of Hematopoietic Cells., Freshney, et al.. Eds. pp. 265-268, Wiley-Liss, Inc., New York, NY. 1994; Hirayama et al., Proc. Natl. Acad. Sci. USA 89:5907-5911, 1992; McNiece and Briddell, in Culture of Hematopoietic Cells, supra; Neben et al., Exp. Hematol. 22:353-359, 1994; Ploemacher and Cobblestone In

Culture of Hematopoietic Cells, supral-21, Spooncer et al, in Culture of Hematopoietic Cells, supral63-179 and Sutherland in Culture of Hematopoietic Cells, supra. 139-162.

Those proteins which exhibit hematopoiesis regulatory activity may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of hematopoeisis is beneficial, such as in the treatment of myeloid or lymphoid cell deficiencies. Involvement in regulating hematopoiesis is indicated even by marginal biological activity in support of colony forming cells or of factor-dependent cell lines. For example, proteins supporting the growth and proliferation of erythroid progenitor cells alone or in combination with other cytokines, indicates utility, for example, in treating various anemias or for use in conjunction with irradiation/chemotherapy to stimulate the production of erythroid precursors and/or erythroid cells. Proteins supporting the growth and proliferation of myeloid cells such as granulocytes and monocytes/macrophages (i.e., traditional CSF activity) may be useful, for example, in conjunction with chemotherapy to prevent or treat consequent myelosuppression. Proteins supporting the growth and proliferation of megakaryocytes and consequently of platelets allows prevention or treatment of various platelet disorders such as thrombocytopenia, and generally may be used in place of or complementary to platelet transfusions. Proteins supporting the growth and proliferation of hematopoietic stem cells which are capable of maturing to any and all of the above-mentioned hematopoietic cells may therefore find therapeutic utility in various stem cell disorders (such as those usually treated with transplantion, including, without limitation, aplastic anemia and paroxysmal nocturnal hemoglobinuria), as well as in repopulating the stem cell compartment post irradiation/chemotherapy, either in vivo or ex vivo (i.e., in conjunction with bone marrow transplantation or with peripheral progenitor cell transplantation (homologous or heterologous)) as normal cells or genetically manipulated for gene therapy. Alternatively, as described in more detail below, genes encoding hematopoiesis regulating activity proteins or nucleic acids regulating the expression of such proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

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#### **EXAMPLE 35**

# Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Regulation of Tissue Growth

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for their effect on tissue growth. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in International Patent Publication No. WO95/16035, International Patent Publication No. WO95/05846 and International Patent Publication No. WO91/07491, which are incorporated herein by reference.

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Assays for wound healing activity include, without limitation, those described in: Winter, *Epidermal Wound Healing*, pps. 71-112, Maibach and Rovee, eds., Year Book Medical Publishers, Inc., Chicago, as modified by Eaglstein and Mertz, *J. Invest. Dermatol.* 71:382-84, 1978, which are incorporated herein by reference.

Those proteins which are involved in the regulation of tissue growth may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of tissue growth is beneficial. For example, a protein encoded by extended cDNAs derived from the 5' ESTs of the present invention also may have utility in compositions used for bone, cartilage, tendon, ligament and/or nerve tissue growth or regeneration, as well as for wound healing and tissue repair and replacement, and in the treatment of burns, incisions and ulcers.

A protein encoded by extended cDNAs derived from the 5' ESTs of the present invention, which induces cartilage and/or bone growth in circumstances where bone is not normally formed, has application in the healing of bone fractures and cartilage damage or defects in humans and other animals. Such a preparation employing a protein of the invention may have prophylactic use in closed as well as open fracture reduction and also in the improved fixation of artificial joints. *De novo* bone synthesis induced by an osteogenic agent contributes to the repair of congenital, trauma induced, or oncologic resection induced craniofacial defects, and also is useful in cosmetic plastic surgery.

A protein of this invention may also be used in the treatment of periodontal disease, and in other tooth repair processes. Such agents may provide an environment to attract bone-forming cells, stimulate growth of bone-forming cells or induce differentiation of bone-forming cell progenitors. A protein of the invention may also be useful in the treatment of osteoporosis or osteoarthritis, such as through stimulation of bone and/or cartilage repair or

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by blocking inflammation or processes of tissue destruction (collagenase activity, osteoclast activity, etc.) mediated by inflammatory processes.

Another category of tissue regeneration activity that may be attributable to the protein encoded by extended cDNAs derived from the 5' ESTs of the present invention is tendon/ligament formation. A protein encoded by extended cDNAs derived from the 5' ESTs of the present invention, which induces tendon/ligament-like tissue or other tissue formation in circumstances where such tissue is not normally formed, has application in the healing of tendon or ligament tears, deformities and other tendon or ligament defects in humans and other animals. Such a preparation employing a tendon/ligament-like tissue inducing protein may have prophylactic use in preventing damage to tendon or ligament tissue, as well as use in the improved fixation of tendon or ligament to bone or other tissues, and in repairing defects to tendon or ligament tissue. De novo tendon/ligament-like tissue formation induced by a composition encoded by extended cDNAs derived from the 5' ESTs of the present invention contributes to the repair of tendon or ligaments defects of congenital, traumatic or other origin and is also useful in cosmetic plastic surgery for attachment or repair of tendons or ligaments. The compositions encoded by extended cDNAs derived from the 5' ESTs of the present invention may provide an environment to attract tendon- or ligamentforming cells, stimulate growth of tendon- or ligament-forming cells, induce differentiation of progenitors of tendon- or ligament-forming cells, or induce growth of tendon/ligament cells or progenitors ex vivo for return in vivo to effect tissue repair. The compositions of the invention may also be useful in the treatment of tendinitis, carpal tunnel syndrome and other tendon or ligament defects. The compositions may also include an appropriate matrix and/or sequestering agent as a carrier as is well known in the art.

The protein encoded by extended cDNAs derived from the 5' ESTs of the present invention may also be useful for proliferation of neural cells and for regeneration of nerve and brain tissue, i.e., for the treatment of central and peripheral nervous system diseases and neuropathies, as well as mechanical and traumatic disorders, which involve degeneration, death or trauma to neural cells or nerve tissue. More specifically, a protein may be used in the treatment of diseases of the peripheral nervous system, such as peripheral nerve injuries, peripheral neuropathy and localized neuropathies, and central nervous system diseases, such as Alzheimer's, Parkinson's disease, Huntington's disease, amyotrophic lateral sclerosis, and

Shy-Drager syndrome. Further conditions which may be treated in accordance with the present invention include mechanical and traumatic disorders, such as spinal cord disorders, head trauma and cerebrovascular diseases such as stroke. Peripheral neuropathies resulting from chemotherapy or other medical therapies may also be treatable using a protein of the invention.

Proteins of the invention may also be useful to promote better or faster closure of non-healing wounds, including without limitation pressure ulcers, ulcers associated with vascular insufficiency, surgical and traumatic wounds, and the like.

It is expected that a protein encoded by extended cDNAs derived from the 5' ESTs of the present invention may also exhibit activity for generation or regeneration of other tissues, such as organs (including, for example, pancreas, liver, intestine, kidney, skin, endothelium) muscle (smooth, skeletal or cardiac) and vascular (including vascular endothelium) tissue, or for promoting the growth of cells comprising such tissues. Part of the desired effects may be by inhibition or modulation of fibrotic scarring to allow normal tissue to generate. A protein of the invention may also exhibit angiogenic activity.

A protein encoded by extended cDNAs derived from the 5' ESTs of the present invention may also be useful for gut protection or regeneration and treatment of lung or liver fibrosis, reperfusion injury in various tissues, and conditions resulting from systemic cytokinc damage.

A protein encoded by extended cDNAs derived from the 5' ESTs of the present invention may also be useful for promoting or inhibiting differentiation of tissues described above from precursor tissues or cells; or for inhibiting the growth of tissues described above.

Alternatively, as described in more detail below, genes encoding tissue growth regulating activity proteins or nucleic acids regulating the expression of such proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

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#### **EXAMPLE 36**

# Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Regulation of Reproductive Hormones

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for their ability to regulate reproductive hormones, such as follicle stimulating hormone. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Vale et al., Endocrinol. 91:562-572, 1972; Ling et al., Nature 321:779-782, 1986; Vale et al., Nature 321:776-779, 1986; Mason et al., Nature 318:659-663, 1985; Forage et al., Proc. Natl. Acad. Sci. USA 83:3091-3095, 1986, Chapter 6.12 in Current Protocols in Immunology, Coligan et al. Eds. Greene Publishing Associates and Wiley-Intersciece; Taub et al., J. Clin. Invest. 95:1370-1376, 1995; Lind et al., APMIS 103:140-146, 1995; Muller et al., Eur. J. Immunol. 25:1744-1748; Gruber et al., J. Immunol. 152:5860-5867, 1994; Johnston et al., J Immunol. 153:1762-1768, 1994.

Those proteins which exhibit activity as reproductive hormones or regulators of cell movement may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of reproductive hormones are beneficial. For example, a protein encoded by extended cDNAs derived from the 5' ESTs of the present invention may also exhibit activinor inhibin-related activities. Inhibins are characterized by their ability to inhibit the release of follicle stimulating hormone (FSH), while activins are characterized by their ability to stimulate the release of FSH. Thus, a protein encoded by extended cDNAs derived from the 5' ESTs of the present invention, alone or in heterodimers with a member of the inhibin a family, may be useful as a contraceptive based on the ability of inhibins to decrease fertility in female mammals and decrease spermatogenesis in male mammals. Administration of sufficient amounts of other inhibins can induce infertility in these mammals. Alternatively, the protein of the invention, as a homodimer or as a heterodimer with other protein subunits of the inhibin-B group, may be useful as a fertility inducing therapeutic, based upon the ability of activin molecules in stimulating FSH release from cells of the anterior pituitary. See, for example, United States Patent 4,798,885, the disclosure of which is incorporated herein by reference. A protein of the invention may also be useful for advancement of the onset of

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fertility in sexually immature mammals, so as to increase the lifetime reproductive performance of domestic animals such as cows, sheep and pigs.

Alternatively, as described in more detail below, genes encoding reproductive hormone regulating activity proteins or nucleic acids regulating the expression of such proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

### **EXAMPLE 37**

# Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Chemotactic/Chemokinetic Activity

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for chemotactic/chemokinetic activity. For example, a protein encoded by extended cDNAs derived from the 5' ESTs of the present invention may have chemotactic or chemokinetic activity (e.g., act as a chemokine) for mammalian cells, including, for example, monocytes, fibroblasts, neutrophils, T-cells, mast cells, eosinophils, epithelial and/or endothelial cells. Chemotactic and chemokinetic proteins can be used to mobilize or attract a desired cell population to a desired site of action. Chemotactic or chemokinetic proteins provide particular advantages in treatment of wounds and other trauma to tissues, as well as in treatment of localized infections. For example, attraction of lymphocytes, monocytes or neutrophils to tumors or sites of infection may result in improved immune responses against the tumor or infecting agent.

A protein or peptide has chemotactic activity for a particular cell population if it can stimulate, directly or indirectly, the directed orientation or movement of such cell population. Preferably, the protein or peptide has the ability to directly stimulate directed movement of cells. Whether a particular protein has chemotactic activity for a population of cells can be readily determined by employing such protein or peptide in any known assay for cell chemotaxis.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Assays for chemotactic activity (which will identify proteins that induce or prevent chemotaxis) consist of assays that measure the ability of a protein to induce the migration of

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cells across a membrane as well as the ability of a protein to induce the adhesion of one cell population to another cell population. Suitable assays for movement and adhesion include, without limitation, those described in: Current Protocols in Immunology, Ed by Coligan, Kruisbeek, Margulies, Shevach and Strober, Pub. Greene Publishing Associates and Wiley-Interscience, Chapter 6.12: 6.12.1-6.12.28; Taub et al., J. Clin. Invest. 95:1370-1376, 1995; Lind et al., APMIS 103:140-146, 1995; Mueller et al., Eur. J. Immunol. 25:1744-1748; Gruber et al., J. Immunol. 152:5860-5867, 1994; Johnston et al. J. Immunol., 153:1762-1768, 1994.

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### **EXAMPLE 38**

# Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Regulation of Blood Clotting

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for their effects on blood clotting. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Linet et al., J. Clin. Pharmacol. 26:131-140, 1986; Burdick et al., Thrombosis Res. 45:413-419, 1987; Humphrey et al., Fibrinolysis 5:71-79, 1991; Schaub, Prostaglandins 35:467-474, 1988.

Those proteins which are involved in the regulation of blood clotting may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of blood clotting is beneficial. For example, a protein of the invention may also exhibit hemostatic or thrombolytic activity. As a result, such a protein is expected to be useful in treatment of various coagulations disorders (including hereditary disorders, such as hemophilias) or to enhance coagulation and other hemostatic events in treating wounds resulting from trauma, surgery or other causes. A protein of the invention may also be useful for dissolving or inhibiting formation of thromboses and for treatment and prevention of conditions resulting therefrom (such as infarction of cardiac and central nervous system vessels (e.g., stroke)). Alternatively, as described in more detail below, genes encoding blood clotting activity proteins or nucleic acids regulating the expression of such proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

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#### **EXAMPLE 39**

# Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Involvement in Receptor/Ligand Interactions

The proteins encoded by the extended cDNAs or a portion thereof may also be evaluated for their involvement in receptor/ligand interactions. Numerous assays for such involvement are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Chapter 7. 7.28.1-7.28.22 in Current Protocols in Immunology, Coligan et al. Eds. Greene Publishing Associates and Wiley-Interscience, Takai et al., Proc. Natl. Acad. Sci. USA 84:6864-6868, 1987; Bierer et al., J. Exp. Med. 168:1145-1156, 1988; Rosenstein et al., J. Exp. Med. 169:149-160, 1989; Stoltenborg et al., J. Immunol. Methods 175:59-68, 1994; Stitt et al., Cell 80:661-670, 1995; Gyuris et al., Cell 75:791-803, 1993.

For example, the proteins encoded by extended cDNAs derived from the 5' ESTs of the present invention may also demonstrate activity as receptors, receptor ligands or inhibitors or agonists of receptor/ligand interactions. Examples of such receptors and ligands include, without limitation, cytokine receptors and their ligands, receptor kinases and their ligands, receptor phosphatases and their ligands, receptors involved in cell-cell interactions and their ligands (including without limitation, cellular adhesion molecules (such as selectins, integrins and their ligands) and receptor/ligand pairs involved in antigen presentation, antigen recognition and development of cellular and humoral immune responses). Receptors and ligands are also useful for screening of potential peptide or small molecule inhibitors of the relevant receptor/ligand interaction. A protein encoded by extended cDNAs derived from the 5' ESTs of the present invention (including, without limitation, fragments of receptors and ligands) may themselves be useful as inhibitors of receptor/ligand interactions. Alternatively, as described in more detail below, genes encoding proteins involved in receptor/ligand interactions or nucleic acids regulating the expression of such proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

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#### **EXAMPLE 40**

# Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Anti-Inflammatory Activity

The proteins encoded by the extended cDNAs or a portion thereof may also be evaluated for anti-inflammatory activity. The anti-inflammatory activity may be achieved by providing a stimulus to cells involved in the inflammatory response, by inhibiting or promoting cell-cell interactions (such as, for example, cell adhesion), by inhibiting or promoting chemotaxis of cells involved in the inflammatory process, inhibiting or promoting cell extravasation, or by stimulating or suppressing production of other factors which more directly inhibit or promote an inflammatory response. Proteins exhibiting such activities can be used to treat inflammatory conditions including chronic or acute conditions, including without limitation inflammation associated with infection (such as septic shock, sepsis or systemic inflammatory response syndrome), ischemia-reperfusioninury, endotoxin lethality, arthritis, complement-mediated hyperacute rejection, nephritis, cytokine- or chemokineinduced lung injury, inflammatory bowel disease, Crohn's disease or resulting from over production of cytokines such as TNF or IL-1. Proteins of the invention may also be useful to treat anaphylaxis and hypersensitivity to an antigenic substance or material. Alternatively, as described in more detail below, genes encoding anti-inflammatory activity proteins or nucleic acids regulating the expression of such proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

#### **EXAMPLE 41**

# Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Tumor Inhibition Activity

The proteins encoded-by the extended cDNAs or a portion thereof may also be evaluated for tumor inhibition activity. In addition to the activities described above for immunological treatment or prevention of tumors, a protein of the invention may exhibit other anti-tumor activities. A protein may inhibit tumor growth directly or indirectly (such as, for example, via ADCC). A protein may exhibit its tumor inhibitory activity by acting on tumor tissue or tumor precursor tissue, by inhibiting formation of tissues necessary to support tumor growth (such as, for example, by inhibiting angiogenesis), by causing production of other

factors, agents or cell types which inhibit tumor growth, or by suppressing, eliminating or inhibiting factors, agents or cell types which promote tumor growth. Alternatively, as described in more detail below, genes tumor inhibition activity proteins or nucleic acids regulating the expression of such proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

A protein of the invention may also exhibit one or more of the following additional activities or effects: inhibiting the growth, infection or function of, or killing, infectious agents, including, without limitation, bacteria, viruses, fungi and other parasites; effecting (suppressing or enhancing) bodily characteristics, including, without limitation, height, weight, hair color, eye color, skin, fat to lean ratio or other tissue pigmentation, or organ or body part size or shape (such as, for example, breast augmentation or diminution, change in bone form or shape); effecting biorhythms or circadian cycles or rhythms; effecting the fertility of male or female subjects; effecting the metabolism, catabolism, anabolism, processing, utilization, storage or elimination of dietary fat, lipid, protein, carbohydrate, vitamins, minerals, cofactors or other nutritional factors or component(s); effecting behavioral characteristics, including, without limitation, appetite, libido, stress, cognition (including cognitive disorders), depression (including depressive disorders) and violent behaviors; providing analgesic effects or other pain reducing effects; promoting differentiation and growth of embryonic stem cells in lineages other than hematopoietic lineages; hormonal or endocrine activity; in the case of enzymes, correcting deficiencies of the enzyme and treating deficiency-related diseases; treatment of hyperproliferative disorders (such as, for example, psoriasis), immunoglobulin-like activity (such as, for example, the ability to bind antigens or complement); and the ability to act as an antigen in a vaccine composition to raise an immune response against such protein or another material or entity which is cross-reactive with such protein. Alternatively, as described in more detail below, genes encoding proteins involved in any of the above mentioned activities or nucleic acids regulating the expression of such proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

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#### **EXAMPLE 42**

# Identification of Proteins which Interact with Polypeptides Encoded by Extended cDNAs

Proteins which interact with the polypeptides encoded by cDNAs derived from the 5' ESTs or fragments thereof, such as receptor proteins, may be identified using two hybrid systems such as the Matchmaker Two Hybrid System 2 (Catalog No. K1604-1, Clontech). As described in the manual accompanying the kit which is incorporated herein by reference, the the cDNAs derived from 5' ESTs, or fragments thereof, are inserted into an expression vector such that they are in frame with DNA encoding the DNA binding domain of the yeast transcriptional activator GAL4. cDNAs in a cDNA library which encode proteins which might interact with the polypeptides encoded by the extended cDNAs or portions thereof are inserted into a second expression vector such that they are in frame with DNA encoding the activation domain of GALA. The two expression plasmids are transformed into yeast and the yeast are plated on selection medium which selects for expression of selectable markers on each of the expression vectors as well as GAL4 dependent expression of the HIS3 gene. Transformants capable of growing on medium lacking histidine are screened for GAL4 dependent lacZ expression. Those cells which are positive in both the histidine selection and the lacZ assay contain plasmids encoding proteins which interact with the polypeptide encoded by the extended cDNAs or portions thereof.

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Alternatively, the system described in Lustig et al., Methods in Enzymology 283: 83-99, 1997, and in U.S. Patent No. 5,654,150, the disclosure of which is incorporated herein by reference, may be used for identifying molecules which interact with the polypeptides encoded by extended cDNAs. In such systems, in vitro transcription reactions are performed on a pool of vectors containing extended cDNA inserts cloned downstream of a promoter which drives in vitro transcription. The resulting pools of mRNAs are introduced into Xenopus laevis oocytes. The oocytes are then assayed for a desired activity.

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Alternatively, the pooled *in vitro* transcription products produced as described above may be translated *in vitro*. The pooled *in vitro* translation products can be assayed for a desired activity or for interaction with a known polypeptide.

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Proteins or other molecules interacting with polypeptides encoded by extended cDNAs can be found by a variety of additional techniques. In one method, affinity

columns containing the polypeptide encoded by the extended cDNA or a portion thereof can be constructed. In some versions, of this method the affinity column contains chimeric proteins in which the protein encoded by the extended cDNA or a portion thereof is fused to glutathione S-transferase. A mixture of cellular proteins or pool of expressed proteins as described above and is applied to the affinity column. Proteins interacting with the polypeptide attached to the column can then be isolated and analyzed on 2-D electrophoresis gel as described in Ramunsen et al., Electrophoresis 18:588-598, 1997, the disclosure of which is incorporated herein by reference. Alternatively, the proteins retained on the affinity column can be purified by electrophoresis based methods and sequenced. The same method can be used to isolate antibodies, to screen phage display products, or to screen phage display human antibodies.

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Proteins interacting with polypeptides encoded by extended cDNAs or portions thereof can also be screened by using an Optical Biosensor as described in Edwards and Leatherbarrow, Analytical Biochemistry 246:1-6, 1997, the disclosure of which is incorporated herein by reference. The main advantage of the method is that it allows the determination of the association rate between the protein and other interacting molecules. Thus, it is possible to specifically select interacting molecules with a high or low association rate. Typically a target molecule is linked to the sensor surface (through a carboxymethl dextran matrix) and a sample of test molecules is placed in contact with the target molecules. The binding of a test molecule to the target molecule causes a change in the refractive index and/ or thickness. This change is detected by the Biosensor provided it occurs in the evanescent field (which extend a few hundred nanometers from the sensor surface). In these screening assays, the target molecule can be one of the polypeptides encoded by extended cDNAs or a portion thereof and the test sample can be a collection of proteins extracted from tissues or cells, a pool of expressed proteins, combinatorial peptide and/ or chemical libraries, or phage displayed peptides. The tissues or cells from which the test proteins are extracted can originate from any species.

In other methods, a target protein is immobilized and the test population is a collection of unique polypeptides encoded by the extended cDNAs or portions thereof.

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To study the interaction of the proteins encoded by the extended cDNAs or portions thereof with drugs, the microdialysis coupled to HPLC method described by Wang et al., Chromatographia 44:205-208, 1997 or the affinity capillary electrophoresis method described by Busch et al., J. Chromatogr. 777:311-328, 1997, the disclosures of which are incorporated herein by reference can be used.

It will be appreciated by those skilled in the art that the proteins expressed from the extended cDNAs or portions may be assayed for numerous activities in addition to those specifically enumerated above. For example, the expressed proteins may be evaluated for applications involving control and regulation of inflammation, tumor proliferation or metastasis, infection, or other clinical conditions. In addition, the proteins expressed from the extended cDNAs or portions thereof may be useful as nutritional agents or cosmetic agents.

The proteins expressed from the cDNAs or portions thereof may be used to generate antibodies capable of specifically binding to the expressed protein or fragments thereof as described in Example 40 below. The antibodies may capable of binding a full length protein encoded by a cDNA derived from a 5' EST, a mature protein (*i.e.* the protein generated by cleavage of the signal peptide) encoded by a cDNA derived from a 5' EST, or a signal peptide encoded by a cDNA derived from a 5' EST. Alternatively, the antibodies may be capable of binding fragments of at least 10 amino acids of the proteins encoded by the above cDNAs. In some embodiments, the antibodies may be capable of binding fragments of at least 15 amino acids of the proteins encoded by the above cDNAs. In other embodiments, the antibodies may be capable of binding fragments of at least 25 amino acids of the proteins expressed from the extended cDNAs which comprise at least 25 amino acids of the proteins encoded by the above cDNAs. In further embodiments, the antibodies may be capable of binding fragments of at least 40 amino acids of the proteins encoded by the above cDNAs.

### **EXAMPLE 43**

## Production of an Antibody to a Human Protein

Substantially pure protein or polypeptide is isolated from the transfected or transformed cells as described in Example 30. The concentration of protein in the final preparation is adjusted, for example, by concentration on an Amicon filter device, to the

level of a few µg/ml. Monoclonal or polyclonal antibody to the protein can then be prepared as follows:

# 1. Monoclonal Antibody Production by Hybridoma Fusion

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Monoclonal antibody to epitopes of any of the peptides identified and isolated as described can be prepared from murine hybridomas according to the classical method of Kohler, and Milstein, Nature 256:495, 1975 or derivative methods thereof. Briefly, a mouse is repetitively inoculated with a few micrograms of the selected protein or peptides derived therefrom over a period of a few weeks. The mouse is then sacrificed, and the antibody producing cells of the spleen isolated. The spleen cells are fused by means of polyethylene glycol with mouse myeloma cells, and the excess unfused cells destroyed by growth of the system on selective media comprising aminopterin (HAT media). The successfully fused cells are diluted and aliquots of the dilution placed in wells of a microtiter plate where growth of the culture is continued. Antibody-producing clones are identified by detection of antibody in the supernatant fluid of the wells by immunoassay procedures, such as ELISA, as originally described by Engvall, Meth. Enzymol. 70:419, 1980, the disclosure of which is incorporated herein by reference and derivative methods thereof. Selected positive clones can be expanded and their monoclonal antibody product harvested for use. Detailed procedures for monoclonal antibody production are described in Davis et al. in Basic Methods in Molecular Biology Elsevier, New York. Section 21-2, the disclosure of which is incorporated herein by reference.

2. Polyclonal Antibody Production by Immunization

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Polyclonal antiserum containing antibodies to heterogenous epitopes of a single protein can be prepared by immunizing suitable animals with the expressed protein or peptides derived therefrom, which can be unmodified or modified to enhance immunogenicity. Effective polyclonal antibody production is affected by many factors related both to the antigen and the host species. For example, small molecules tend to be less immunogenic than others and may require the use of carriers and adjuvant. Also, host animals response vary depending on site of inoculations and doses, with both inadequate or

excessive doses of antigen resulting in low titer antisera. Small doses (ng level) of antigen administered at multiple intradermal sites appears to be most reliable. An effective immunization protocol for rabbits can be found in Vaitukaitis. *et al*, *J. Clin. Endocrinol. Metab.* 33:988-991 (1971), the disclosure of which is incorporated herein by reference.

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Booster injections can be given at regular intervals, and antiserum harvested when antibody titer thereof, as determined semi-quantitatively, for example, by double immunodiffusion in agar against known concentrations of the antigen, begins to fall. See, for example, Ouchterlony, et al., Chap. 19 in: Handbook of Experimental Immunology D. Wier (ed) Blackwell (1973), the disclosure of which is incorporated herein by reference. Plateau concentration of antibody is usually in the range of 0.1 to 0.2 mg/ml of serum (about 12 µM). Affinity of the antisera for the antigen is determined by preparing competitive binding curves, as described, for example, by Fisher, D., Chap. 42 in: Manual of Clinical Immunology, 2d Ed. (Rose and Friedman, Eds.) Amer. Soc. For Microbiol., Washington, D.C. (1980), the disclosure of which is incorporated herein by reference..

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Antibody preparations prepared according to either protocol are useful in quantitative immunoassays which determine concentrations of antigen-bearing substances in biological samples; they are also used semi-quantitatively or qualitatively to identify the presence of antigen in a biological sample. The antibodies may also be used in therapeutic compositions for killing cells expressing the protein or reducing the levels of the protein in the body.

# V. Use of 5' ESTs or Sequences Obtainable Therefrom or Portions Thereof as Reagents

The 5' ESTs of the present invention (or cDNAs or genomic DNAs obtainable therefrom) may be used as reagents in isolation procedures, diagnostic assays, and forensic procedures. For example, sequences from the 5' ESTs (or cDNAs or genomic DNAs obtainable therefrom) may be detectably labeled and used as probes to isolate other sequences capable of hybridizing to them. In addition, sequences from 5' ESTs (or cDNAs or genomic DNAs obtainable therefrom) may be used to design PCR primers to be used in isolation, diagnostic, or forensic procedures.

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# 1. Use of 5' ESTs or Sequences Obtainable Therefrom or Portions Thereof in Isolation, Diagnostic and Forensic Procedures

### **EXAMPLE 44**

# Preparation of PCR Primers and Amplification of DNA

The 5' EST sequences (or cDNAs or genomic DNAs obtainable therefrom) may be used to prepare PCR primers for a variety of applications, including isolation procedures for cloning nucleic acids capable of hybridizing to such sequences, diagnostic techniques and forensic techniques. The PCR primers are at least 10 bases, and preferably at least 12, 15, or 17 bases in length. More preferably, the PCR primers are at least 20-30 bases in length. In some embodiments, the PCR primers may be more than 30 bases in length. It is preferred that the primer pairs have approximately the same G/C ratio, so that melting temperatures are approximately the same. A variety of PCR techniques are familiar to those skilled in the art. For a review of PCR technology, see Molecular Cloning to Genetic Engineering, White Ed. in Methods in Molecular Biology 67: Humana Press, Totowa 1997, the disclosure of which is incorporated herein by reference. In each of these PCR procedures, PCR primers on either side of the nucleic acid sequences to be amplified are added to a suitably prepared nucleic acid sample along with dNTPs and a thermostable polymerase such as Taq polymerase, Pfu polymerase, or Vent polymerase. The nucleic acid in the sample is denatured and the PCR primers are specifically hybridized to complementary nucleic acid sequences in the sample. The hybridized primers are extended. Thereafter, another cycle of denaturation, hybridization, and extension is initiated. The cycles are repeated multiple times to produce an amplified fragment containing the nucleic acid sequence between the primer sites.

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### **EXAMPLE 45**

## Use of 5'ESTs as Probes

Probes derived from 5' ESTs (or cDNAs or genomic DNAs obtainable therefrom), including full length cDNAs or genomic sequences, may be labeled with detectable labels familiar to those skilled in the art, including radioisotopes and non-radioactive labels, to provide a detectable probe. The detectable probe may be single stranded or double stranded and may be made using techniques known in the art, including *in vitro* transcription, nick

translation, or kinase reactions. A nucleic acid sample containing a sequence capable of hybridizing to the labeled probe is contacted with the labeled probe. If the nucleic acid in the sample is double stranded, it may be denatured prior to contacting the probe. In some applications, the nucleic acid sample may be immobilized on a surface such as a nitrocellulose or nylon membrane. The nucleic acid sample may comprise nucleic acids obtained from a variety of sources, including genomic DNA, cDNA libraries, RNA, or tissue samples.

Procedures used to detect the presence of nucleic acids capable of hybridizing to the detectable probe include well known techniques such as Southern blotting, Northern blotting, dot blotting, colony hybridization, and plaque hybridization. In some applications, the nucleic acid capable of hybridizing to the labeled probe may be cloned into vectors such as expression vectors, sequencing vectors, or *in vitro* transcription vectors to facilitate the characterization and expression of the hybridizing nucleic acids in the sample. For example, such techniques may be used to isolate and clone sequences in a genomic library or cDNA library which are capable of hybridizing to the detectable probe as described in Example 30 above.

PCR primers made as described in Example 44 above may be used in forensic analyses, such as the DNA fingerprinting techniques described in Examples 46-50 below. Such analyses may utilize detectable probes or primers based on the sequences of the the 5' ESTs or of cDNAs or genomic DNAs isolated using the 5' ESTs.

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#### **EXAMPLE 46**

## Forensic Matching by DNA Sequencing

In one exemplary method, DNA samples are isolated from forensic specimens of, for example, hair, semen, blood or skin cells by conventional methods. A panel of PCR primers based on a number of the 5' ESTs of Example 25, or cDNAs or genomic DNAs isolated therefrom as described above, is then utilized in accordance with Example 44 to amplify DNA of approximately 100-200 bases in length from the forensic specimen. Corresponding sequences are obtained from a test subject. Each of these identification DNAs is then sequenced using standard techniques, and a simple database comparison determines the differences, if any, between the sequences from the subject and those from the sample. Statistically significant differences between the suspect's DNA sequences and those from the sample conclusively prove a lack of identity. This lack of identity can be proven, for example,

with only one sequence. Identity, on the other hand, should be demonstrated with a large number of sequences, all matching. Preferably, a minimum of 50 statistically identical sequences of 100 bases in length are used to prove identity between the suspect and the sample.

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## **EXAMPLE 47**

# Positive Identification by DNA Sequencing

The technique outlined in the previous example may also be used on a larger scale to provide a unique fingerprint-type identification of any individual. In this technique, primers are prepared from a large number of 5'EST sequences from Example 25, or cDNA or genomic DNA sequences obtainable therefrom. Preferably, 20 to 50 different primers are used. These primers are used to obtain a corresponding number of PCR-generated DNA segments from the individual in question in accordance with Example 44. Each of these DNA segments is sequenced, using the methods set forth in Example 46. The database of sequences generated through this procedure uniquely identifies the individual from whom the sequences were obtained. The same panel of primers may then be used at any later time to absolutely correlate tissue or other biological specimen with that individual.

### **EXAMPLE 48**

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### Southern Blot Forensic Identification

The procedure of Example 47 is repeated to obtain a panel of at least 10 amplified sequences from an individual and a specimen. Preferably, the panel contains at least 50 amplified sequences. More preferably, the panel contains 100 amplified sequences. In some embodiments, the panel contains 200 amplified sequences. This PCR-generated DNA is then digested with one or a combination of, preferably, four base specific restriction enzymes. Such enzymes are commercially available and known to those of skill in the art. After digestion, the resultant gene fragments are size separated in multiple duplicate wells on an agarose gel and transferred to nitrocellulose using Southern blotting techniques well known to those with skill in the art. For a review of Southern blotting see Davis *et al.* (Basic Methods in Molecular Biology, 1986, Elsevier Press. pp 62-65), the disclosure of which is incorporated herein by reference.

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A panel of probes based on the sequences of 5' ESTs (or cDNAs or genomic DNAs obtainable therefrom), or fragments thereof of at least 10 bases, are radioactively or colorimetrically labeled using methods known in the art, such as nick translation or end labeling, and hybridized to the Southern blot using techniques known in the art (Davis *et al.*, supra). Preferably, the probe comprises at least 12, 15, or 17 consecutive nucleotides from the 5' EST (or cDNAs or genomic DNAs obtainable therefrom). More preferably, the probe comprises at least 20-30 consecutive nucleotides from the 5' EST (or cDNAs or genomic DNAs obtainable therefrom). In some embodiments, the probe comprises more than 30 nucleotides from the 5' EST (or cDNAs or genomic DNAs obtainable therefrom).

Preferably, at least 5 to 10 of these labeled probes are used, and more preferably at least about 20 or 30 are used to provide a unique pattern. The resultant bands appearing from the hybridization of a large sample of 5' EST (or cDNAs or genomic DNAs obtainable therefrom) will be a unique identifier. Since the restriction enzyme cleavage will be different for every individual, the band pattern on the Southern blot will also be unique. Increasing the number of 5' EST (or cDNAs or genomic DNAs obtainable therefrom) probes will provide a statistically higher level of confidence in the identification since there will be an increased number of sets of bands used for identification.

#### **EXAMPLE 49**

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## **Dot Blot Identification Procedure**

Another technique for identifying individuals using the 5' EST sequences disclosed herein utilizes a dot blot hybridization technique.

Genomic DNA is isolated from nuclei of subject to be identified. Oligonucleotide probes of approximately 30 bp in length are synthesized that correspond to at least 10, preferably 50 sequences from the 5' ESTs or cDNAs or genomic DNAs obtainable therefrom. The probes are used to hybridize to the genomic DNA through conditions known to those in the art. The oligonucleotides are end labeled with P<sup>32</sup> using polynucleotide kinase (Pharmacia). Dot Blots are created by spotting the genomic DNA onto nitrocellulose or the like using a vacuum dot blot manifold (BioRad, Richmond California). The nitrocellulose filter containing the genomic sequences is baked or UV linked to the filter, prehybridized and hybridized with labeled probe using techniques known in the art (Davis et al., supra). The <sup>32</sup>P

labeled DNA fragments are sequentially hybridized with successively stringent conditions to detect minimal differences between the 30 bp sequence and the DNA. Tetramethylammonium chloride is useful for identifying clones containing small numbers of nucleotide mismatches (Wood et al., Proc. Natl. Acad. Sci. USA 82(6):1585-1588, 1985) which is hereby incorporated by reference. A unique pattern of dots distinguishes one individual from another individual.

5' EST sequences (or cDNAs or genomic DNAs obtainable therefrom) or oligonucleotides containing at least 10 consecutive bases from these sequences can be used as probes in the following alternative fingerprinting technique. Preferably, the probe comprises at least 12, 15, or 17 consecutive nucleotides from the 5' EST sequences (or cDNAs or genomic DNAs obtainable therefrom). More preferably, the probe comprises at least 20-30 consecutive nucleotides from the 5' EST sequences (or cDNAs or genomic DNAs obtainable therefrom). In some embodiments, the probe comprises more than 30 nucleotides from the 5' EST sequences (or cDNAs or genomic DNAs obtainable therefrom).

Preferably, a plurality of probes having sequences from different genes are used in the alternative fingerprinting technique. Example 50 below provides a representative alternative fingerprinting procedure in which the probes are derived from 5'EST.

#### **EXAMPLE 50**

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### Alternative "Fingerprint" Identification Technique

20-mer oligonucleotides are prepared from a large number, e.g. 50, 100, or 200, of 5'EST using commercially available oligonucleotide services such as Genset, Paris, France. Cell samples from the test subject are processed for DNA using techniques well known to those with skill in the art. The nucleic acid is digested with restriction enzymes such as EcoRI and XbaI. Following digestion, samples are applied to wells for electrophoresis. The procedure, as known in the art, may be modified to accommodate polyacrylamide electrophoresis, however in this example, samples containing 5 ug of DNA are loaded into wells and separated on 0.8% agarose gels. The gels are transferred onto nitrocellulose using standard Southern blotting techniques.

10 ng of each of the oligonucleotides are pooled and end-labeled with <sup>32</sup>P. The nitrocellulose is prehybridized with blocking solution and hybridized with the labeled probes.

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Following hybridization and washing, the nitrocellulose filter is exposed to X-Omat AR X-ray film. The resulting hybridization pattern will be unique for each individual.

It is additionally contemplated within this example that the number of probe sequences used can be varied for additional accuracy or clarity.

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The proteins encoded by the extended cDNAs may also be used to generate antibodies as explained in Examples 30 and 43 in order to identify the tissue type or cell species from which a sample is derived as described in example 51.

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#### **EXAMPLE 51**

# Identification of Tissue Types or Cell Species by Means of Labeled Tissue Specific Antibodies

Identification of specific tissues is accomplished by the visualization of tissue specific antigens by means of antibody preparations according to Examples 30 and 43 which are conjugated, directly or indirectly to a detectable marker. Selected labeled antibody species bind to their specific antigen binding partner in tissue sections, cell suspensions, or in extracts of soluble proteins from a tissue sample to provide a pattern for qualitative or semi-qualitative interpretation.

Antisera for these procedures must have a potency exceeding that of the native preparation, and for that reason, antibodies are concentrated to a mg/ml level by isolation of the gamma globulin fraction, for example, by ion-exchange chromatography or by ammonium sulfate fractionation. Also, to provide the most specific antisera, unwanted antibodies, for example to common proteins, must be removed from the gamma globulin fraction, for example by means of insoluble immunoabsorbents, before the antibodies are labeled with the marker. Either monoclonal or heterologous antisera is suitable for either procedure.

#### A. Immunohistochemical techniques

Purified, high-titer antibodies, prepared as described above, are conjugated to a detectable marker, as described, for example, by Fudenberg, Chap. 26 in: Basic and Clinical Immunology, 3rd Ed. Lange, Los Altos, California, 1980, or Rose, et al., Chap. 12 in: Methods in Immunodiagnosis, 2d Ed. John Wiley and Sons, New York (1980), the disclosures of which are incorporated herein by reference.

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A fluorescent marker, either fluorescein or rhodamine, is preferred, but antibodies can also be labeled with an enzyme that supports a color producing reaction with a substrate, such as horseradish peroxidase. Markers can be added to tissue-bound antibody in a second step, as described below. Alternatively, the specific antitissue antibodies can be labeled with ferritin or other electron dense particles, and localization of the ferritin coupled antigen-antibody complexes achieved by means of an electron microscope. In yet another approach, the antibodies are radiolabeled, with, for example <sup>125</sup>I, and detected by overlaying the antibody treated preparation with photographic emulsion.

Preparations to carry out the procedures can comprise monoclonal or polyclonal antibodies to a single protein or peptide identified as specific to a tissue type, for example, brain tissue, or antibody preparations to several antigenically distinct tissue specific antigens can be used in panels, independently or in mixtures, as required.

Tissue sections and cell suspensions are prepared for immunohistochemical examination according to common histological techniques. Multiple cryostat sections (about 4 µm, unfixed) of the unknown tissue and known control, are mounted and each slide covered with different dilutions of the antibody preparation. Sections of known and unknown tissues should also be treated with preparations to provide a positive control, a negative control, for example, pre-immune sera, and a control for non-specific staining, for example, buffer.

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Treated sections are incubated in a humid chamber for 30 min at room temperature, rinsed, then washed in buffer for 30-45 min. Excess fluid is blotted away, and the marker developed.

If the tissue specific antibody was not labeled in the first incubation, it can be labeled at this time in a second antibody-antibody reaction, for example, by adding fluorescein- or enzyme-conjugated antibody against the immunoglobulin class of the antiserum-producing species, for example, fluorescein labeled antibody to mouse IgG. Such labeled sera are commercially available.

The antigen found in the tissues by the above procedure can be quantified by measuring the intensity of color or fluorescence on the tissue section, and calibrating that signal using appropriate standards.

## B. Identification of tissue specific soluble proteins

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The visualization of tissue specific proteins and identification of unknown tissues from that procedure is carried out using the labeled antibody reagents and detection strategy as described for immunohistochemistry; however the sample is prepared according to an electrophoretic technique to distribute the proteins extracted from the tissue in an orderly array on the basis of molecular weight for detection.

A tissue sample is homogenized using a Virtis apparatus; cell suspensions are disrupted by Dounce homogenization or osmotic lysis, using detergents in either case as required to disrupt cell membranes, as is the practice in the art. Insoluble cell components such as nuclei, microsomes, and membrane fragments are removed by ultracentrifugation, and the soluble protein-containing fraction concentrated if necessary and reserved for analysis.

A sample of the soluble protein solution is resolved into individual protein species by conventional SDS polyacrylamide electrophoresis as described, for example, by Davis, et al., Section 19-2 in: Basic Methods in Molecular Biology, Leder ed., Elsevier, New York, 1986. the disclosure of which is incorporated herein by reference, using a range of amounts of polyacrylamide in a set of gels to resolve the entire molecular weight range of proteins to be detected in the sample. A size marker is run in parallel for purposes of estimating molecular weights of the constituent proteins. Sample size for analysis is a convenient volume of from 5 to 55 µl, and containing from about 1 to 100 µg protein. An aliquot of each of the resolved proteins is transferred by blotting to a nitrocellulose filter paper, a process that maintains the pattern of resolution. Multiple copies are prepared. The procedure, known as Western Blot Analysis, is well described in Davis, L. et al., supra Section 19-3. One set of nitrocellulose blots is stained with Coomassie blue dye to visualize the entire set of proteins for comparison with the antibody bound proteins. The remaining nitrocellulose filters are then incubated with a solution of one or more specific antisera to tissue specific proteins prepared as described in Examples 30 and 43. In this procedure, as in procedure A above, appropriate positive and negative sample and reagent controls are run.

In either procedure A or B, a detectable label can be attached to the primary tissue antigen-primary antibody complex according to various strategies and permutations thereof. In a straightforward approach, the primary specific antibody can be labeled; alternatively, the unlabeled complex can be bound by a labeled secondary anti-IgG antibody. In other

approaches, either the primary or secondary antibody is conjugated to a biotin molecule, which can, in a subsequent step, bind an avidin conjugated marker. According to yet another strategy, enzyme labeled or radioactive protein A, which has the property of binding to any IgG, is bound in a final step to either the primary or secondary antibody.

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The visualization of tissue specific antigen binding at levels above those seen in control tissues to one or more tissue specific antibodies, prepared from the gene sequences identified from extended cDNA sequences, can identify tissues of unknown origin, for example, forensic samples, or differentiated tumor tissue that has metastasized to foreign bodily sites.

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In addition to their applications in forensics and identification, 5' ESTs (or cDNAs or genomic DNAs obtainable therefrom) may be mapped to their chromosomal locations. Example 52 below describes radiation hybrid (RH) mapping of human chromosomal regions using 5'ESTs. Example 53 below describes a representative procedure for mapping an 5' EST to its location on a human chromosome. Example 54 below describes mapping of 5' ESTs on metaphase chromosomes by Fluorescence In Situ Hybridization (FISH). Those skilled in the art will appreciate that the method of Examples 52-54 may also be used to map cDNAs or genomic DNAs obtainable from the 5' ESTs to their chromosomal locations.

# 2. Use of 5' ESTs or Sequences Obtainable Therefrom or Portions Thereof in Chromosome Mapping

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### **EXAMPLE 52**

# Radiation hybrid mapping of 5'ESTs to the human genome

Radiation hybrid (RH) mapping is a somatic cell genetic approach that can be used for high resolution mapping of the human genome. In this approach, cell lines containing one or more human chromosomes are lethally irradiated, breaking each chromosome into fragments whose size depends on the radiation dose. These fragments are rescued by fusion with cultured rodent cells, yielding subclones containing different portions of the human genome. This technique is described by Benham et al., Genomics 4:509-517, 1989; and Cox et al., Science 250:245-250, 1990, the entire contents of which are hereby incorporated by reference. The random and independent nature of the subclones permits efficient mapping of

any human genome marker. Human DNA isolated from a panel of 80-100 cell lines provides a mapping reagent for ordering 5'EST. In this approach, the frequency of breakage between markers is used to measure distance, allowing construction of fine resolution maps as has been done using conventional ESTs (Schuler *et al.*, *Science* 274:540-546, 1996, hereby incorporated by reference).

RH mapping has been used to generate a high-resolution whole genome radiation hybrid map of human chromosome 17q22-q25.3 across the genes for growth hormone (GH) and thymidine kinase (TK) (Foster et al., Genomics 33:185-192, 1996), the region surrounding the Gorlin syndrome gene (Obermayr et al., Eur. J. Hum. Genet. 4:242-245, 1996), 60 loci covering the entire short arm of chromosome 12 (Raeymaekers et al., Genomics 29:170-178, 1995), the region of human chromosome 22 containing the neurofibromatosis type 2 locus (Frazer et al., Genomics 14:574-584, 1992) and 13 loci on the long arm of chromosome 5 (Warrington et al., Genomics 11:701-708, 1991).

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#### **EXAMPLE 53**

# Mapping of 5'ESTs to HumanChromosomes using PCR techniques

5' ESTs (or cDNAs or genomic DNAs obtainable therefrom) may be assigned to human chromosomes using PCR based methodologies. In such approaches, oligonucleotide primer pairs are designed from the 5' ESTs (or cDNAs or genomic DNAs obtainable therefrom) to minimize the chance of amplifying through an intron. Preferably, the oligonucleotide primers are 18-23 bp in length and are designed for PCR amplification. The creation of PCR primers from known sequences is well known to those with skill in the art. For a review of PCR technology see Erlich in PCR Technology, Principles and Applications for DNA Amplification, Freeman and Co., New York, 1992, the disclosure of which is incorporated herein by reference.

The primers are used in polymerase chain reactions (PCR) to amplify templates from total human genomic DNA. PCR conditions are as follows: 60 ng of genomic DNA is used as a template for PCR with 80 ng of each oligonucleotide primer, 0.6 unit of Taq polymerase, and 1 μCu of a <sup>32</sup>P-labeled deoxycytidine triphosphate. The PCR is performed in a microplate thermocycler (Techne) under the following conditions: 30 cycles of 94°C, 1.4 min; 55°C, 2 min; and 72°C, 2 min; with a final extension at 72°C for 10 min. The amplified

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products are analyzed on a 6% polyacrylamide sequencing gel and visualized by autoradiography. If the length of the resulting PCR product is identical to the distance between the ends of the primer sequences in the extended cDNA from which the primers are derived, then the PCR reaction is repeated with DNA templates from two panels of human-rodent somatic cell hybrids, BIOS PCRable DNA (BIOS Corporation) and NIGMS Human-Rodent Somatic Cell Hybrid Mapping Panel Number 1 (NIGMS, Camden, NJ).

PCR is used to screen a series of somatic cell hybrid cell lines containing defined sets of human chromosomes for the presence of a given 5' EST (or cDNA or genomic DNA obtainable therefrom). DNA is isolated from the somatic hybrids and used as starting templates for PCR\_reactions using the primer pairs from the 5' EST (or cDNA or genomic DNA obtainable therefrom). Only those somatic cell hybrids with chromosomes containing the human gene corresponding to the 5' EST (or cDNA or genomic DNA obtainable therefrom) will yield an amplified fragment. The 5' EST (or cDNA or genomic DNA obtainable therefrom) are assigned to a chromosome by analysis of the segregation pattern of PCR products from the somatic hybrid DNA templates. The single human chromosome present in all cell hybrids that give rise to an amplified fragment is the chromosome containing that 5'EST (or cDNA or genomic DNA obtainable therefrom). For a review of techniques and analysis of results from somatic cell gene mapping experiments, see Ledbetter et al., Genomics 6:475-481, 1990, the disclosure of which is incorporated herein by reference.

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## **EXAMPLE 54**

# Mapping of Extended 5' ESTs to Chromosomes Using Fluorescence In Situ Hybridization

Fluorescence in situ hybridization allows the 5'EST (or cDNA or genomic DNA obtainable therefrom) to be mapped to a particular location on a given chromosome. The chromosomes to be used for fluorescence in situ hybridization techniques may be obtained from a variety of sources including cell cultures, tissues, or whole blood.

In a preferred embodiment, chromosomal localization of an 5'EST (or cDNA or genomic DNA obtainable therefrom) is obtained by FISH as described by Cherif et al. (*Proc. Natl. Acad. Sci. U.S.A.*, 87:6639-6643, 1990), the disclosure of which is incorporated herein by reference. Metaphase chromosomes are prepared from phytohemagglutinin (PHA)-

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stimulated blood cell donors. PHA-stimulated lymphocytes from healthy males are cultured for 72 h in RPMI-1640 medium. For synchronization, methotrexate (10 µM) is added for 17 h, followed by addition of 5-bromodeoxyuridine (5-BrdU, 0.1 mM) for 6 h. Colcemid (1 µg/ml) is added for the last 15 min before harvesting the cells. Cells are collected, washed in RPMI, incubated with a hypotonic solution of KCl (75 mM) at 37°C for 15 min and fixed in three changes of methanol:acetic acid (3:1). The cell suspension is dropped onto a glass slide and air dried. The 5'EST (or cDNA or genomic DNA obtainable therefrom) is labeled with biotin-16 dUTP by nick translation according to the manufacturer's instructions (Bethesda Research Laboratories, Bethesda, MD), purified using a Sephadex G-50 column (Pharmacia, Upsala, Sweden) and precipitated. Just prior to hybridization, the DNA pellet is dissolved in hybridization buffer (50% formamide, 2 X SSC, 10% dextran sulfate, 1 mg/ml sonicated salmon sperm DNA, pH 7) and the probe is denatured at 70°C for 5-10 min.

Slides kept at -20°C are treated for 1 h at 37°C with RNase A (100 µg/ml), rinsed three times in 2 X SSC and dehydrated in an ethanol series. Chromosome preparations are denatured in 70% formamide, 2 X SSC for 2 min at 70°C, then dehydrated at 4°C. The slides are treated with proteinase K (10 µg/100 ml in 20 mM Tris-HCl, 2 mM CaCl<sub>2</sub>) at 37°C for 8 min and dehydrated. The hybridization mixture containing the probe is placed on the slide, covered with a coverslip, sealed with rubber cement and incubated overnight in a humid chamber at 37°C. After hybridization and post-hybridization washes, the biotinylated probe is detected by avidin-FITC and amplified with additional layers of biotinylated goat anti-avidin and avidin-FITC. For chromosomal localization, fluorescent R-bands are obtained as previously described (Cherif et al., supra.). The slides are observed under a LEICA fluorescence microscope (DMRXA). Chromosomes are counterstained with propidium iodide and the fluorescent signal of the probe appears as two symmetrical yellow-green spots on both chromatids of the fluorescent R-band chromosome (red). Thus, a particular 5'EST (or cDNA or genomic DNA obtainable therefrom) may be localized to a particular cytogenetic R-band on a given chromosome.

Once the 5'EST (or cDNA or genomic DNA obtainable therefrom) have been assigned to particular chromosomes using the techniques described in Examples 52-54 above,

they may be utilized to construct a high resolution map of the chromosomes on which they are located or to identify the chromosomes in a sample.

#### **EXAMPLE 55**

## Use of 5'EST to Construct or Expand Chromosome Maps

Chromosome mapping involves assigning a given unique sequence to a particular chromosome as described above. Once the unique sequence has been mapped to a given chromosome, it is ordered relative to other unique sequences located on the same chromosome. One approach to chromosome mapping utilizes a series of yeast artificial chromosomes (YACs) bearing several thousand long inserts derived from the chromosomes of the organism from which the extended cDNAs (or genomic DNAs obtainable therefrom) are obtained. This approach is described in Nagaraja et al., Genome Research 7:210-222. 1997, the disclosure of which is incorporated herein by reference. Briefly, in this approach each chromosome is broken into overlapping pieces which are inserted into the YAC vector. The YAC inserts are screened using PCR or other methods to determine whether they include the 5'EST (or cDNA or genomic DNA obtainable therefrom) whose position is to be determined. Once an insert has been found which includes the 5'EST (or cDNA or genomic DNA obtainable therefrom), the insert can be analyzed by PCR or other methods to determine whether the insert also contains other sequences known to be on the chromosome or in the region from which the 5'EST (or cDNA or genomic DNA obtainable therefrom) was derived. This process can be repeated for each insert in the YAC library to determine the location of each of the extended cDNAs (or genomic DNAs obtainable therefrom) relative to one another and to other known chromosomal markers. In this way, a high resolution map of the distribution of numerous unique markers along each of the organisms chromosomes may be obtained.

As described in Example 56 below extended cDNAs (or genomic DNAs obtainable therefrom) may also be used to identify genes associated with a particular phenotype, such as hereditary disease or drug response.

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# 3. Use of 5'ESTs or Sequences Obtained Therefrom or Fragments Thereof in Gene Identification

#### **EXAMPLE 56**

# Identification of genes associated with hereditary diseases or drug response

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This example illustrates an approach useful for the association of 5'ESTs (or cDNA or genomic DNA obtainable therefrom) with particular phenotypic characteristics. In this example, a particular 5'EST (or cDNA or genomic DNA obtainable therefrom) is used as a test probe to associate that 5'EST (or cDNA or genomic DNA obtainable therefrom) with a particular phenotypic characteristic.

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5'ESTs (or cDNA or genomic DNA obtainable therefrom) are mapped to a particular location on a human chromosome using techniques such as those described in Examples 52 and 53 or other techniques known in the art. A search of Mendelian Inheritance in Man (McKusick in *Mendelian Inheritance in Man* (available on line through Johns Hopkins University Welch Medical Library) reveals the region of the human chromosome which contains the 5'EST (or cDNA or genomic DNA obtainable therefrom) to be a very gene rich region containing several known genes and several diseases or phenotypes for which genes have not been identified. The gene corresponding to this 5'EST (or cDNA or genomic DNA obtainable therefrom) thus becomes an immediate candidate for each of these genetic diseases.

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Cells from patients with these diseases or phenotypes are isolated and expanded in culture. PCR primers from the 5'EST (or cDNA or genomic DNA obtainable therefrom) are used to screen genomic DNA, mRNA or cDNA obtained from the patients. 5'ESTs (or cDNA or genomic DNA obtainable therefrom) that are not amplified in the patients can be positively associated with a particular disease by further analysis. Alternatively, the PCR analysis may yield fragments of different lengths when the samples are derived from an individual having the phenotype associated with the disease than when the sample is derived from a healthy individual, indicating that the gene containing the 5'EST may be responsible for the genetic disease.

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# VI. Use of 5'EST (or cDNA or Genomic DNA Obtainable Therefrom) to Construct Vectors

The present 5'ESTs (or cDNA or genomic DNA obtainable therefrom) may also be used to construct secretion vectors capable of directing the secretion of the proteins encoded by genes therein. Such secretion vectors may facilitate the purification or enrichment of the proteins encoded by genes inserted therein by reducing the number of background proteins from which the desired protein must be purified or enriched. Exemplary secretion vectors are described in Example 57 below.

### 1. Construction of Secretion Vectors

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### **EXAMPLE 57**

#### Construction of Secretion Vectors

The secretion vectors include a promoter capable of directing gene expression in the host cell, tissue, or organism of interest. Such promoters include the Rous Sarcoma Virus promoter, the SV40 promoter, the human cytomegalovirus promoter, and other promoters familiar to those skilled in the art.

A signal sequence from a 5' EST (or cDNAs or genomic DNAs obtainable therefrom) is operably linked to the promoter such that the mRNA transcribed from the promoter will direct the translation of the signal peptide. The host cell, tissue, or organism may be any cell, tissue, or organism which recognizes the signal peptide encoded by the signal sequence in the 5' EST (or cDNA or genomic DNA obtainable therefrom). Suitable hosts include mammalian cells, tissues or organisms, avian cells, tissues, or organisms, insect cells, tissues or organisms, or yeast.

In addition, the secretion vector contains cloning sites for inserting genes encoding the proteins which are to be secreted. The cloning sites facilitate the cloning of the insert gene in frame with the signal sequence such that a fusion protein in which the signal peptide is fused to the protein encoded by the inserted gene is expressed from the mRNA transcribed from the promoter. The signal peptide directs the extracellular secretion of the fusion protein.

The secretion vector may be DNA or RNA and may integrate into the chromosome of the host, be stably maintained as an extrachromosomal replicon in the host, be an artificial chromosome, or be transiently present in the host. Many nucleic acid backbones suitable for

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use as secretion vectors are known to those skilled in the art, including retroviral vectors, SV40 vectors, Bovine Papilloma Virus vectors, yeast integrating plasmids, yeast episomal plasmids, yeast artificial chromosomes, human artificial chromosomes, P element vectors, baculovirus vectors, or bacterial plasmids capable of being transiently introduced into the host.

The secretion vector may also contain a polyA signal such that the polyA signal is located downstream of the gene inserted into the secretion vector.

After the gene encoding the protein for which secretion is desired is inserted into the secretion vector, the secretion vector is introduced into the host cell, tissue, or organism using calcium phosphate precipitation, DEAE-Dextran, electroporation, liposome-mediated transfection, viral particles or as naked DNA. The protein encoded by the inserted gene is then purified or enriched from the supernatant using conventional techniques such as ammonium sulfate precipitation, immunoprecipitation, immunochromatography, size exclusion chromatography, ion exchange chromatography, and HPLC. Alternatively, the secreted protein may be in a sufficiently enriched or pure state in the supernatant or growth media of the host to permit it to be used for its intended purpose without further enrichment.

The signal sequences may also be inserted into vectors designed for gene therapy. In such vectors, the signal sequence is operably linked to a promoter such that mRNA transcribed from the promoter encodes the signal peptide. A cloning site is located downstream of the signal sequence such that a gene encoding a protein whose secretion is desired may readily be inserted into the vector and fused to the signal sequence. The vector is introduced into an appropriate host cell. The protein expressed from the promoter is secreted extracellularly, thereby producing a therapeutic effect.

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The 5' ESTs may also be used to clone sequences located upstream of the 5' ESTs which are capable of regulating gene expression, including promoter sequences, enhancer sequences, and other upstream sequences which influence transcription or translation levels. Once identified and cloned, these upstream regulatory sequences may be used in expression vectors designed to direct the expression of an inserted gene in a desired spatial, temporal, developmental, or quantitative fashion. Example 58 describes a method for cloning sequences upstream of the extended cDNAs or 5' ESTs.

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# 2. Identification of Upstream Sequences With Promoting or Regulatory Activities

### **EXAMPLE 58**

Use of Extended cDNAs or 5' ESTs to Clone Upstream Sequences from Genomic DNA

Sequences derived from extended cDNAs or 5' ESTs may be used to isolate the promoters of the corresponding genes using chromosome walking techniques. In one chromosome walking technique, which utilizes the GenomeWalker<sup>TM</sup> kit available from Clontech, five complete genomic DNA samples are each digested with a different restriction enzyme which has a 6 base recognition site and leaves a blunt end. Following digestion, oligonucleotide adapters are ligated to each end of the resulting genomic DNA fragments.

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For each of the five genomic DNA libraries, a first PCR reaction is performed according to the manufacturer's instructions (which are incorporated herein by reference) using an outer adaptor primer provided in the kit and an outer gene specific primer. The gene specific primer should be selected to be specific for the extended cDNA or 5' EST of interest and should have a melting temperature, length, and location in the extended cDNA or 5'EST which is consistent with its use in PCR reactions. Each first PCR reaction contains 5 ng of genomic DNA, 5 µl of 10X Tth reaction buffer, 0.2 mM of each dNTP, 0.2 µM each of outer adaptor primer and outer gene specific primer, 1.1 mM of Mg(OAc)<sub>2</sub>, and 1 µl of the Tth polymerase 50X mix in a total volume of 50 µl. The reaction cycle for the first PCR reaction is as follows: 1 min - 94°C / 2 sec - 94°C, 3 min - 72°C (7 cycles) / 2 sec - 94°C, 3 min - 67°C.

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The product of the first PCR reaction is diluted and used as a template for a second PCR reaction according to the manufacturer's instructions using a pair of nested primers which are located internally on the amplicon resulting from the first PCR reaction. For example, 5 µl of the reaction product of the first PCR reaction mixture may be diluted 180 times. Reactions are made in a 50 µl volume having a composition identical to that of the first PCR reaction except the nested primers are used. The first nested primer is specific for the adaptor, and is provided with the GenomeWalker<sup>TM</sup> kit. The second nested primer is specific for the particular extended cDNA or 5' EST for which the promoter is to be cloned and should have a melting temperature, length, and location in the extended cDNA or 5' EST which is consistent with its use in PCR

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reactions. The reaction parameters of the second PCR reaction are as follows: 1 min - 94°C / 2 sec - 94°C, 3 min - 72°C (6 cycles) / 2 sec - 94°C, 3 min - 67°C (25 cycles) / 5 min - 67°C. The product of the second PCR reaction is purified, cloned, and sequenced using standard techniques.

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Alternatively, two or more human genomic DNA libraries can be constructed by using two or more restriction enzymes. The digested genomic DNA is cloned into vectors which can be converted into single stranded, circular, or linear DNA. A biotinylated oligonucleotide comprising at least 15 nucleotides from the extended cDNA or 5' EST sequence is hybridized to the single stranded DNA. Hybrids between the biotinylated oligonucleotide and the single stranded DNA containing the extended cDNA or EST sequence are isolated as described in Example 29 above. Thereafter, the single stranded DNA containing the extended cDNA or EST sequence is released from the beads and converted into double stranded DNA using a primer specific for the extended cDNA or 5' EST sequence or a primer corresponding to a sequence included in the cloning vector. The resulting double stranded DNA is transformed into bacteria. DNAs containing the 5' EST or extended cDNA sequences are identified by colony PCR or colony hybridization.

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Once the upstream genomic sequences have been cloned and sequenced as described above, prospective promoters and transcription start sites within the upstream sequences may be identified by comparing the sequences upstream of the extended cDNAs or 5' ESTs with databases containing known transcription start sites, transcription factor binding sites, or promoter sequences.

In addition, promoters in the upstream sequences may be identified using promoter reporter vectors as described in Example.

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### **EXAMPLE 59**

# Identification of Promoters in Cloned Upstream Sequences

The genomic sequences upstream of the extended cDNAs or 5' ESTs are cloned into a suitable promoter reporter vector, such as the pSEAP-Basic, pSEAP-Enhancer, pβgal-Basic, pβgal-Enhancer, or pEGFP-1 Promoter Reporter vectors available from Clontech. Briefly, each of these promoter reporter vectors include multiple cloning sites positioned

upstream of a reporter gene encoding a readily assayable protein such as secreted alkaline phosphatase, β galactosidase, or green fluorescent protein. The sequences upstream of the extended cDNAs or 5′ ESTs are inserted into the cloning sites upstream of the reporter gene in both orientations and introduced into an appropriate host cell. The level of reporter protein is assayed and compared to the level obtained from a vector which lacks an insert in the cloning site. The presence of an elevated expression level in the vector containing the insert with respect to the control vector indicates the presence of a promoter in the insert. If necessary, the upstream sequences can be cloned into vectors which contain an enhancer for augmenting transcription levels from weak promoter sequences. A significant level of expression above that observed with the vector lacking an insert indicates that a promoter sequence is present in the inserted upstream sequence.

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Appropriate host cells for the promoter reporter vectors may be chosen based on the results of the above described determination of expression patterns of the extended cDNAs and ESTs. For example, if the expression pattern analysis indicates that the mRNA corresponding to a particular extended cDNA or 5' EST is expressed in fibroblasts, the promoter reporter vector may be introduced into a human fibroblast cell line.

Promoter sequences within the upstream genomic DNA may be further defined by constructing nested deletions in the upstream DNA using conventional techniques such as Exonuclease III digestion. The resulting deletion fragments can be inserted into the promoter reporter vector to determine whether the deletion has reduced or obliterated promoter activity. In this way, the boundaries of the promoters may be defined. If desired, potential individual regulatory sites within the promoter may be identified using site directed mutagenesis or linker scanning to obliterate potential transcription factor binding sites within the promoter individually or in combination. The effects of these mutations on transcription levels may be determined by inserting the mutations into the cloning sites in the promoter reporter vectors.

#### **EXAMPLE 60**

# Cloning and Identification of Promoters

Using the method described in Example 58 above with 5' ESTs, sequences upstream of several genes were obtained. Using the primer pairs GGG AAG ATG GAG ATA GTA

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TTG CCT G (SEQ ID NO:29) and CTG CCA TGT ACA TGA TAG AGA GAT TC (SEQ ID NO:30), the promoter having the internal designation P13H2 (SEQ ID NO:31) was obtained.

Using the primer pairs GTA CCA GGGG ACT GTG ACC ATT GC (SEQ ID NO:32) and CTG TGA CCA TTG CTC CCA AGA GAG (SEQ ID NO:33), the promoter having the internal designation P15B4 (SEQ ID NO:34) was obtained.

Using the primer pairs CTG GGA TGG AAG GCA CGG TA (SEQ ID NO:35) and GAG ACC ACA CAG CTA GAC AA (SEQ ID NO:36), the promoter having the internal designation P29B6 (SEQ ID NO:37) was obtained.

Figure 4 provides a schematic description of the promoters isolated and the way they are assembled with the corresponding 5' tags. The upstream sequences were screened for the presence of motifs resembling transcription factor binding sites or known transcription start sites using the computer program MatInspector release 2.0, August 1996.

Table VII describes the transcription factor binding sites present in each of these promoters. The columns labeled matrice provides the name of the MatInspector matrix used. The column labeled position provides the 5' position of the promoter site. Numeration of the sequence starts from the transcription site as determined by matching the genomic sequence with the 5' EST sequence. The column labeled "orientation" indicates the DNA strand on which the site is found, with the + strand being the coding strand as determined by matching the genomic sequence with the sequence of the 5' EST. The column labeled "score" provides the MatInspector score found for this site. The column labeled "length" provides the length of the site in nucleotides. The column labeled "sequence" provides the sequence of the site found.

Bacterial clones containing plasmids containing the promoter sequences described above described above are presently stored in the inventor's laboratories under the internal identification numbers provided above. The inserts may be recovered from the deposited materials by growing an aliquot of the appropriate bacterial clone in the appropriate medium. The plasmid DNA can then be isolated using plasmid isolation procedures familiar to those skilled in the art such as alkaline lysis minipreps or large scale alkaline lysis plasmid isolation procedures. If desired the plasmid DNA may be further enriched by centrifugation on a cesium chloride gradient, size exclusion chromatography, or anion exchange chromatography.

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The plasmid DNA obtained using these procedures may then be manipulated using standard cloning techniques familiar to those skilled in the art. Alternatively, a PCR can be done with primers designed at both ends of the EST insertion. The PCR product which corresponds to the 5' EST can then be manipulated using standard cloning techniques familiar to those skilled in the art.

The promoters and other regulatory sequences located upstream of the extended cDNAs or 5' ESTs may be used to design expression vectors capable of directing the expression of an inserted gene in a desired spatial, temporal, developmental, or quantitative manner. A promoter capable of directing the desired spatial, temporal, developmental, and quantitative patterns may be selected using the results of the expression analysis described in Example 26 above. For example, if a promoter which confers a high level of expression in muscle is desired, the promoter sequence upstream of an extended cDNA or 5' EST derived from an mRNA which is expressed at a high level in muscle, as determined by the method of Example 26, may be used in the expression vector.

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Preferably, the desired promoter is placed near multiple restriction sites to facilitate the cloning of the desired insert downstream of the promoter, such that the promoter is able to drive expression of the inserted gene. The promoter may be inserted in conventional nucleic acid backbones designed for extrachromosomal replication, integration into the host chromosomes or transient expression. Suitable backbones for the present expression vectors include retroviral backbones, backbones from eukaryotic episomes such as SV40 or Bovine Papilloma Virus, backbones from bacterial episomes, or artificial chromosomes.

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Preferably, the expression vectors also include a polyA signal downstream of the multiple restriction sites for directing the polyadenylation of mRNA transcribed from the gene inserted into the expression vector.

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Following the identification of promoter sequences using the procedures of Examples 58-60, proteins which interact with the promoter may be identified as described in Example 61 below.

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#### **EXAMPLE 61**

## Identification of Proteins Which Interact with Promoter Sequences, Upstream Regulatory Sequences, or mRNA

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Sequences within the promoter region which are likely to bind transcription factors may be identified by homology to known transcription factor binding sites or through conventional mutagenesis or deletion analyses of reporter plasmids containing the promoter sequence. For example, deletions may be made in a reporter plasmid containing the promoter sequence of interest operably linked to an assayable reporter gene. The reporter plasmids carrying various deletions within the promoter region are transfected into an appropriate host cell and the effects of the deletions on expression levels is assessed. Transcription factor binding sites within the regions in which deletions reduce expression levels may be further localized using site directed mutagenesis, linker scanning analysis, or other techniques familiar to those skilled in the art.

Nucleic acids encoding proteins which interact with sequences in the promoter may be identified using one-hybrid systems such as those described in the manual accompanying the Matchmaker One-Hybrid System kit available from Clontech (Catalog No. K1603-1), the disclosure of which is incorporated herein by reference. Briefly, the Matchmaker One-hybrid system is used as follows. The target sequence for which it is desired to identify binding proteins is cloned upstream of a selectable reporter gene and integrated into the yeast genome. Preferably, multiple copies of the target sequences are inserted into the reporter plasmid in tandem. A library comprised of fusions between cDNAs to be evaluated for the ability to bind to the promoter and the activation domain of a yeast transcription factor, such as GALA, is transformed into the yeast strain containing the integrated reporter sequence. The yeast are plated on selective media to select cells expressing the selectable marker linked to the promoter sequence. The colonies which grow on the selective media contain genes encoding proteins which bind the target sequence. The inserts in the genes encoding the fusion proteins are further characterized by sequencing. In addition, the inserts may be inserted into expression vectors or in vitro transcription vectors. Binding of the polypeptides encoded by the inserts to the promoter DNA may be confirmed by techniques familiar to those skilled in the art, such as gel shift analysis or DNAse protection analysis.

## VII. Use of 5' ESTs (or cDNAs or Genomic DNAs Obtainable Therefrom) in Gene Therapy

The present invention also comprises the use of 5'ESTs (or cDNA or genomic DNA obtainable therefrom) in gene therapy strategies, including antisense and triple helix strategies as described in Examples 62 and 63 below. In antisense approaches, nucleic acid sequences complementary to an mRNA are hybridized to the mRNA intracellularly, thereby blocking the expression of the protein encoded by the mRNA. The antisense sequences may prevent gene expression through a variety of mechanisms. For example, the antisense sequences may inhibit the ability of ribosomes to translate the mRNA. Alternatively, the antisense sequences may block transport of the mRNA from the nucleus to the cytoplasm, thereby limiting the amount of mRNA available for translation. Another mechanism through which antisense sequences may inhibit gene expression is by interfering with mRNA splicing. In yet another strategy, the antisense nucleic acid may be incorporated in a ribozyme capable of specifically cleaving the target mRNA.

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#### **EXAMPLE 62**

### Preparation and Use of Antisense Oligonucleotides

The antisense nucleic acid molecules to be used in gene therapy may be either DNA or RNA sequences. They may comprise a sequence complementary to the sequence of the 5'EST (or cDNA or genomic DNA obtainable therefrom). The antisense nucleic acids should have a length and melting temperature sufficient to permit formation of an intracellular duplex with sufficient stability to inhibit the expression of the mRNA in the duplex. Strategies for designing antisense nucleic acids suitable for use in gene therapy are disclosed in Green et al., Ann. Rev. Biochem. 55:569-597, 1986; and Izant and Weintraub, Cell 36:1007-1015, 1984, which are hereby incorporated by reference.

In some strategies, antisense molecules are obtained from a nucleotide sequence encoding a protein by reversing the orientation of the coding region with respect to a promoter so as to transcribe the opposite strand from that which is normally transcribed in the cell. The antisense molecules may be transcribed using *in vitro* transcription systems such as those which employ T7 or SP6 polymerase to generate the transcript. Another approach

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involves transcription of the antisense nucleic acids in vivo by operably linking DNA containing the antisense sequence to a promoter in an expression vector.

Alternatively, oligonucleotides which are complementary to the strand normally transcribed in the cell may be synthesized *in vitro*. Thus, the antisense nucleic acids are complementary to the corresponding mRNA and are capable of hybridizing to the mRNA to create a duplex. In some embodiments, the antisense sequences may contain modified sugar phosphate backbones to increase stability and make them less sensitive to RNase activity. Examples of modifications suitable for use in antisense strategies are described by Rossi *et al.*, *Pharmacol. Ther.* **50(2)**:245-254, 1991, which is hereby incorporated by reference.

Various types of antisense oligonucleotides complementary to the sequence of the 5'EST (or cDNA or genomic DNA obtainable therefrom) may be used. In one preferred embodiment, stable and semi-stable antisense oligonucleotides described in International Application No. PCT WO94/23026, hereby incorporated by reference, are used. In these molecules, the 3' end or both the 3' and 5' ends are engaged in intramolecular hydrogen bonding between complementary base pairs. These molecules are better able to withstand exonuclease attacks and exhibit increased stability compared to conventional antisense oligonucleotides.

In another preferred embodiment, the antisense oligodeoxynucleotides against herpes simplex virus types 1 and 2 described in International Application No. WO 95/04141, hereby incorporated by reference, are used.

In yet another preferred embodiment, the covalently cross-linked antisense oligonucleotides described in International Application No. WO 96/31523, hereby incorporated by reference, are used. These double- or single-stranded oligonucleotides comprise one or more, respectively, inter- or intra-oligonucleotide covalent cross-linkages, wherein the linkage consists of an amide bond between a primary amine group of one strand and a carboxyl group of the other strand or of the same strand, respectively, the primary amine group being directly substituted in the 2' position of the strand nucleotide monosaccharide ring, and the carboxyl group being carried by an aliphatic spacer group substituted on a nucleotide or nucleotide analog of the other strand or the same strand, respectively.

The antisense oligodeoxynucleotides and oligonucleotides disclosed in International Application No. WO 92/18522, incorporated by reference, may also be used. These molecules are stable to degradation and contain at least one transcription control recognition sequence which binds to control proteins and are effective as decoys therefore. These molecules may contain "hairpin" structures, "dumbbell" structures, "modified dumbbell" structures, "cross-linked" decoy structures and "loop" structures.

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In another preferred embodiment, the cyclic double-stranded oligonucleotides described in European Patent Application No. 0 572 287 A2, hereby incorporated by reference are used. These ligated oligonucleotide "dumbbells" contain the binding site for a transcription factor and inhibit expression of the gene under control of the transcription factor by sequestering the factor.

Use of the closed antisense oligonucleotides disclosed in International Application No. WO 92/19732, hereby incorporated by reference, is also contemplated. Because these molecules have no free ends, they are more resistant to degradation by exonucleases than are conventional oligonucleotides... These oligonucleotides may be multifunctional, interacting with several regions which are not adjacent to the target mRNA.

The appropriate level of antisense nucleic acids required to inhibit gene expression may be determined using *in vitro* expression analysis. The antisense molecule may be introduced into the cells by diffusion, injection, infection, transfection or h-region-mediated import using procedures known in the art. For example, the antisense nucleic acids can be introduced into the body as a bare or naked oligonucleotide, oligonucleotide encapsulated in lipid, oligonucleotide sequence encapsidated by viral protein, or as an oligonucleotide operably linked to a promoter contained in an expression vector. The expression vector may be any of a variety of expression vectors known in the art, including retroviral or viral vectors, vectors capable of extrachromosomal replication, or integrating vectors. The vectors may be DNA or RNA.

The antisense molecules are introduced onto cell samples at a number of different concentrations preferably between  $1\times10^{-10}$ M to  $1\times10^{-4}$ M. Once the minimum concentration that can adequately control gene expression is identified, the optimized dose is translated into a dosage suitable for use *in vivo*. For example, an inhibiting concentration in culture of  $1\times10^{-7}$  translates into a dose of approximately 0.6 mg/kg bodyweight. Levels of oligonucleotide

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approaching 100 mg/kg bodyweight or higher may be possible after testing the toxicity of the oligonucleotide in laboratory animals. It is additionally contemplated that cells from the vertebrate are removed, treated with the antisense oligonucleotide, and reintroduced into the vertebrate.

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It is further contemplated that the antisense oligonucleotide sequence is incorporated into a ribozyme sequence to enable the antisense to specifically bind and cleave its target mRNA. For technical applications of ribozyme and antisense oligonucleotides see Rossi et al., supra.

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In a preferred application of this invention, the polypeptide encoded by the gene is first identified, so that the effectiveness of antisense inhibition on translation can be monitored using techniques that include but are not limited to antibody-mediated tests such as RIAs and ELISA, functional assays, or radiolabeling.

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The 5' ESTs of the present invention (or cDNAs or genomic DNAs obtainable therefrom) may also be used in gene therapy approaches based on intracellular triple helix formation. Triple helix oligonucleotides are used to inhibit transcription from a genome. They are particularly useful for studying alterations in cell activity as it is associated with a particular gene. The 5' EST sequences (or cDNAs or genomic DNAs obtainable therefrom) of the present invention or, more preferably, a portion of those sequences, can be used to inhibit gene expression in individuals having diseases associated with expression of a particular gene. Similarly, a portion of 5' EST sequences (or cDNAs or genomic DNAs obtainable therefrom) can be used to study the effect of inhibiting transcription of a particular gene within a cell. Traditionally, homopurine sequences were considered the most useful for triple helix strategies. However, homopyrimidine sequences can also inhibit gene expression. Such homopyrimidine oligonucleotides bind to the major groove at homopurine:homopyrimidine sequences. Thus, both types of sequences from the 5'EST or from the gene corresponding to the 5'EST are contemplated within the scope of this invention.

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#### **EXAMPLE 63**

## Preparation and Use of Triple Helix Probes

The sequences of the 5' ESTs (or cDNAs or genomic DNAs obtainable therefrom) are scanned to identify 10-mer to 20-mer homopyrimidine or homopurine stretches which could be used in triple-helix based strategies for inhibiting gene expression. Following identification of candidate homopyrimidine or homopurine stretches, their efficiency in inhibiting gene expression is assessed by introducing varying amounts of oligonucleotides containing the candidate sequences into tissue culture cells which normally express the target gene. The oligonucleotides may be prepared on an oligonucleotide synthesizer or they may be purchased commercially from a company specializing in custom oligonucleotide synthesis, such as GENSET, Paris, France.

The oligonucleotides may be introduced into the cells using a variety of methods known to those skilled in the art, including but not limited to calcium phosphate precipitation, DEAE-Dextran, electroporation, liposome-mediated transfection or native uptake.

Treated cells are monitored for altered cell function or reduced gene expression using techniques such as Northern blotting, RNase protection assays, or PCR based strategies to monitor the transcription levels of the target gene in cells which have been treated with the oligonucleotide. The cell functions to be monitored are predicted based upon the homologies of the target gene corresponding to the extended cDNA from which the oligonucleotide was

derived with known gene sequences that have been associated with a particular function. The cell functions can also be predicted based on the presence of abnormal physiologies within cells derived from individuals with a particular inherited disease, particularly when the

The oligonucleotides which are effective in inhibiting gene expression in tissue culture cells may then be introduced *in vivo* using the techniques described above and in Example 62

extended cDNA is associated with the disease using techniques described in Example 56.

at a dosage calculated based on the in vitro results, as described in Example 62.

In some embodiments, the natural (beta) anomers of the oligonucleotide units can be replaced with alpha anomers to render the oligonucleotide more resistant to nucleases. Further, an intercalating agent such as ethidium bromide, or the like, can be attached to the 3' end of the alpha oligonucleotide to stabilize the triple helix. For information on the

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generation of oligonucleotides suitable for triple helix formation see Griffin et al., Science 245:967-971, 1989, which is hereby incorporated by this reference.

#### **EXAMPLE 64**

# Use of cDNAs Obtained Using the 5' ESTs to Express an Encoded Protein in a Host Organism

The cDNAs obtained as described above using the 5' ESTs of the present invention may also be used to express an encoded protein in a host organism to produce a beneficial effect. In such procedures, the encoded protein may be transiently expressed in the host organism or stably expressed in the host organism. The encoded protein may have any of the activities described above. The encoded protein may be a protein which the host organism lacks or, alternatively, the encoded protein may augment the existing levels of the protein in the host organism.

A full length extended cDNA encoding the signal peptide and the mature protein, or an extended cDNA encoding only the mature protein is introduced into the host organism. The extended cDNA may be introduced into the host organism using a variety of techniques known to those of skill in the art. For example, the extended cDNA may be injected into the host organism as naked DNA such that the encoded protein is expressed in the host organism, thereby producing a beneficial effect.

Alternatively, the extended cDNA may be cloned into an expression vector downstream of a promoter which is active in the host organism. The expression vector may be any of the expression vectors designed for use in gene therapy, including viral or retroviral vectors. The expression vector may be directly introduced into the host organism such that the encoded protein is expressed in the host organism to produce a beneficial effect. In another approach, the expression vector may be introduced into cells *in vitro*. Cells containing the expression vector are thereafter selected and introduced into the host organism, where they express the encoded protein to produce a beneficial effect.

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#### **EXAMPLE 65**

## Use of Signal Peptides Encoded by 5' ESTs or Sequences obtained Therefrom to Import Proteins Into Cells

The short core hydrophobic region (h) of signal peptides encoded by the 5'ESTS or extended cDNAs derived from SEQ ID NOs: 38-270 may also be used as a carrier to import a peptide or a protein of interest, so-called cargo, into tissue culture cells (Lin et al., J. Biol. Chem., 270: 14225-14258, 1995; Du et al., J. Peptide Res., 51: 235-243, 1998; Rojas et al., Nature Biotech., 16: 370-375, 1998).

When cell permeable peptides of limited size (approximately up to 25 amino acids) are to be translocated across cell membrane, chemical synthesis may be used in order to add the h region to either the C-terminus or the N-terminus to the cargo peptide of interest. Alternatively, when longer peptides or proteins are to be imported into cells, nucleic acids can be genetically engineered, using techniques familiar to those skilled in the art, in order to link the extended cDNA sequence encoding the h region to the 5' or the 3' end of a DNA sequence coding for a cargo polypeptide. Such genetically engineered nucleic acids are then translated either *in vitro* or *in vivo* after transfection into appropriate cells, using conventional techniques to produce the resulting cell permeable polypeptide. Suitable hosts cells are then simply incubated with the cell permeable polypeptide which is then translocated across the membrane.

This method may be applied to study diverse intracellular functions and cellular processes. For instance, it has been used to probe functionally relevant domains of intracellular proteins and to examine protein-protein interactions involved in signal transduction pathways (Lin et al., supra; Lin et al., J. Biol. Chem., 271: 5305-5308, 1996; Rojas et al., J. Biol. Chem., 271: 27456-27461, 1996; Liu et al., Proc. Natl. Acad. Sci. USA, 93: 11819-11824, 1996; Rojas et al., Bioch. Biophys. Res. Commun., 234: 675-680, 1997).

Such techniques may be used in cellular therapy to import proteins producing therapeutic effects. For instance, cells isolated from a patient may be treated with imported therapeutic proteins and then re-introduced into the host organism.

Alternatively, the h region of signal peptides of the present invention could be used in combination with a nuclear localization signal to deliver nucleic acids into cell nucleus. Such oligonucleotides may be antisense oligonucleotides or oligonucleotides designed to form

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triple helixes, as described in examples 62 and 63 respectively, in order to inhibit processing and/or maturation of a target cellular RNA.

As discussed above, the cDNAs or portions thereof obtained using the 5' ESTs of the present invention can be used for various purposes. The polynucleotides can be used to express recombinant protein for analysis, characterization or therapeutic use; as markers for tissues in which the corresponding protein is preferentially expressed (either constitutively or at a particular stage of tissue differentiation or development or in disease states); as molecular weight markers on Southern gels; as chromosome markers or tags (when labeled) to identify chromosomes or to map related gene positions; to compare with endogenous DNA sequences in patients to identify potential genetic disorders, as probes to hybridize and thus discover novel, related DNA sequences; as a source of information to derive PCR primers for genetic fingerprinting; for selecting and making oligomers for attachment to a "gene chip" or other support, including for examination for expression patterns; to raise anti-protein antibodies using DNA immunization techniques; and as an antigen to raise anti-DNA antibodies or elicit another immune response. Where the polynucleotide encodes a protein which binds or potentially binds to another protein (such as, for example, in a receptor-ligand interaction), the polynucleotide can also be used in interaction trap assays (such as, for example, that described in Gyuris et al., Cell 75:791-803, 1993, the disclosure of which is hereby incorporated by reference) to identify polynucleotides encoding the other protein with which binding occurs or to identify inhibitors of the binding interaction.

The proteins or polypeptides provided by the present invention can similarly be used in assays to determine biological activity, including in a panel of multiple proteins for high-throughput screening; to raise antibodies or to elicit another immune response; as a reagent (including the labeled reagent) in assays designed to quantitatively determine levels of the protein (or its receptor) in biological fluids; as markers for tissues in which the corresponding protein is preferentially expressed (either constitutively or at a particular stage of tissue differentiation or development or in a disease state); and, of course, to isolate correlative receptors or ligands. Where the protein binds or potentially binds to another protein (such as, for example, in a receptor-ligand interaction), the protein can be used to identify the other protein with which binding occurs or to identify inhibitors of the binding interaction. Proteins

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involved in these binding interactions can also be used to screen for peptide or small molecule inhibitors or agonists of the binding interaction.

Any or all of these research utilities are capable of being developed into reagent grade or kit format for commercialization as research products.

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Methods for performing the uses listed above are well known to those skilled in the art. References disclosing such methods include without limitation *Molecular Cloning*; A Laboratory Manual, 2d ed., Cold Spring Harbor Laboratory Press, Sambrook, Fritsch and Maniatis eds., 1989, and Methods in Enzymology; Guide to Molecular Cloning Techniques, Academic Press, Berger and Kimmel eds., 1987.

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Polynucleotides and proteins of the present invention can also be used as nutritional sources or supplements. Such uses include without limitation use as a protein or amino acid supplement, use as a carbon source, use as a nitrogen source and use as a source of carbohydrate. In such cases the protein or polynucleotide of the invention can be added to the feed of a particular organism or can be administered as a separate solid or liquid preparation, such as in the form of powder, pills, solutions, suspensions or capsules. In the case of microorganisms, the protein or polynucleotide of the invention can be added to the medium in or on which the microorganism is cultured.

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Although this invention has been described in terms of certain preferred embodiments, other embodiments which will be apparent to those of ordinary skill in the art in view of the disclosure herein are also within the scope of this invention. Accordingly, the scope of the invention is intended to be defined only by reference to the appended claims. All documents cited herein are incorporated herein by reference in their entirety.

	Search characteristic	teristic	Selection	Selection Characteristics	
Step	Program	Strand	Parameters	Identity (%)	Length (bp)
miscellanaeous	blastn	both	S=61 X=16	06	17
tRNA	fasta	both	•	08	90
rRNA	blastn	both	S=108	80	40
mtRNA	blastn	both	S=108	08	40
Procaryotic	blastn	both	S=144	06	40
Fungat	blastn	both	S=144	06	40
Alu	fasta*	both	•	02	40
L1	blastn	both	S=72	0/	40
Repeats	blastn	both	S=72	0.2	40
Promoters	blastn	top	S=54 X=16	06	15†
Vertebrate	fasta*	both	S=108	06	30
ESTs	blastn	both	S=108 X=16	06	30
Proteins	blastx¤	top	E = 0.001	1	•

Table 1: Parameters used for each step of EST analysis

use "Quick Fast" Database scanner alignement further constrained to begin closer than 10bp to EST\5' end using BLOSUM62 substitution matrix

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TABLE II

SEQ. ID		VON HEIJNE	TISSUE	INTERNAL
NO.	CATEGORY	SCORE	SOURCE	DESIGNATION
	<u>0.11.2001.1</u>	<u> </u>	SOURCE	DESIGNATION
ID38	new	10.8	Brain	33-19-2-H2-PU
ID39	new	10.8	Brain	33-56-1-E8-PU
ID40	new	10.0	Brain	33-79-3-D12-PU
ID41	new	9.6		· · · · · - · - · · ·
ID42	new	9.5	Brain	33-72-2-B2-PU
ID43	new	9.1	Brain	33-13-2-B9-PU
ID43	new	9	Brain	33-113-1-E9-PU
ID45	new	8.8	Brain	33-28-4-E8-PU
ID46			Brain	33-12-3-F2-PU
ID47	new	8.8	Brain	33-70-1-C11-PU
ID47 ID48	new	8.5	Brain	33-74-1-B2-PU
	new	8.5	Brain	33-29-3-F1-PU
ID49	new	8.4	Brain	33-8-2-A1-PU
ID50	new	8.3	Brain	17-17-3-A9-PU
ID51	new	8.3	Brain	33-106-2-A8-PU
ID52	new	8.3	Brain	33-112-4-E7-PU
ID53	new	8.2	Brain	33-98-1-E6-PU
ID54	new	8.2	Brain	33-76-1-B6-PU
ID55	new	8	Brain	33-35-4-G8-PU
ID56	new	7.9	Brain	33-17-3-E4-PU
ID57	new	7.9	Brain	33-110-4-B5-PU
ID58	new	7.8	Brain	33-40-1-A11-PU
ID59	new	7.7	Brain	33-71-1-A8-PU
ID60	new	7.7	Brain	33-96-3-G7-PU
ID61	new	7.6	Brain	33-112-3-D12-PU
ID62	new	7.6	Brain	33-62-2-B3-PU
ID63	new	7.6	Brain	33-6-4-G6-PU
ID64	new	7.5	Brain	33-82-4-E2-PU
ID65	new	7.4	Brain	33-81-3-H11-PU
ID66	new	7.3	Brain	33-64-1-B4-PU
ID67	new	7.2	Brain	33-31-1-B12-PU
ID68	new	7	Brain	33-24-4-F9-PU
ID69	new	7	Brain	33-110-3-E9-PU
ID70	new	7	Brain	
ID71	new	6.9		33-4-2-G5-PU
ID72	new	6.9	Brain	33-74-2-A4-PU
ID73		6.9	Brain	33-52-4-F9-PU
ID74	new		Brain	33-74-1-B11-PU
	new	6.8	Brain	33-10-4-D9-PU
ID75	new	6.8	Brain	33-15-2-H3-PU
ID76	new	6.7	Brain	33-38-2-D5-PU
ID77	new	6.7	Brain	33-78-3-D2-PU
ID78	new	6.7	Brain	33-96-3-D3-PU
ID79	new	6.6	Brain	33-76-4-B11-PU
ID80	new	6.3	Brain	33-39-1-C6-PU
ID81	new	6.1	Brain	33-106-3-B12-PU
ID82	new	6	Brain	33-4-2-B7-PU
ID83	new	5.9	Brain	33-99-2-E4-PU
ID84	new	5.9	Brain	33-34-1-B1-PU
ID85	new	5.8	Brain	33-67-4-E9-PU
ID86	new	5.7	Brain	33-11-3-H11-PU

SEQ. ID		VON HEIJNE	TISSUE	INTERNAL
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ID87 ID88	new	5.6	Brain	33-13-2-A8-PU
ID89	new	5.6	Brain	33-83-4-B6-PU
ID99	new	5.6	Brain	33-70-1-E4-PU
ID90 ID91	new	5.6	Brain	33-5-3-H11-PU
ID91 ID92	new	5.6	Brain	33-10-3-G5-PU
ID92 ID93	new	5.5	Brain	33-97-4-G4-PU
ID93 ID94	new	5.5	Brain	33-46-4-F4-PU
ID95	new	5.4	Brain	33-4-1-G11-PU
· ID96	new	5.3	Brain	33-105-1-H5-PU
ID97	new	5.3	Brain	33-74-2-B10-PU
ID98	new	5.3	Brain	33-49-3-E5-PU
ID98	new	5.3	Brain	33-114-2-A1-PU
ID100	new	5.2	Brain	33-71-1-G12-PU
ID101	new	5.2	Brain	33-47-3-E6-PU
ID101 ID102	new	5.2	Brain	33-1-2-E8-PU
ID102	new	5.2	Brain	33-93-4-E12-PU
	new	5.1	Brain	33-1-2-H1-PU
ID104	new	5.1	Brain	17-10-1-H8-PU
ID105	new	5	Brain	33-110-2-B8-PU
ID106	new	5	Brain	33-104-3-D9-PU
ID107	new	5	Brain	33-72-2-H11-PU
ID108	new	4.9	Brain	33-7-4-D6-PU
ID109	new	4.9	Brain	33-31-4-G2-PU
ID110	new	4.9	Brain	33-109-1-E8-PU
ID111	new	4.8	Brain	17-1-2-B11-PU
ID112	new	4.8	Brain	33-19-4-H3-PU
ID113	new	4.8	Brain	33-14-4-E1-PU
ID114	new	4.8	Brain	33-70-3-H1-PU
ID115	new	4.7	Brain	33-86-4-H10-PU
ID116	new .	4.7	Brain	33-107-3-D5-PU
ID117	new	4.7	Brain	33-23-4-B9-PU
ID118	new	4.7	Brain	33-82-4-H5-PU
ID119	new	4.6	Brain	33-16-3-F4-PU
ID120	new	4.6	Brain	33-97-4-C5-PU
ID121	new	4.6	Brain	33-100-3-B10-PU
ID122	new	4.6	Brain	33-59-3-E3-PU
ID123	new	4.5	Brain	33-25-1-G2-PU
ID124	new	4.5	Brain	17-16-3-B2-PU
ID125	new	4.4	Brain	33-52-4-E7-PU
ID126	new	4.4	Brain	33-91-1-D1-PU
ID127	new	4.4	Brain	33-26-1-B9-PU
ID128	new	4.4	Brain	33-97-3-H6-PU
ID129	new	4.4	Brain	33-109-2-E8-PU
ID130	new	4.3	Brain	33-59-2-B7-PU
ID131	new	4.3	Brain	33-28-4-D1-PU
ID132	new	4.3	Brain	33-29-4-E2-PU
ID133	new	4.1	Brain	33-70-1-H6-PU
ID134	new	4.1	Brain	33-7-1-B2-PU
ID135	new	4.1	Brain	33-52-4-F8-PU
ID136	new	4.1	Brain	33-23-2-A6-PU
ID137	new.	4. I	Brain	33-39-3-E5-PU

SEQ. ID		VON HEIJNE	TISSUE	INTERNAL
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ID138	new	4.1	Brain	33-81-4-H6-PU
ID139	new	4.1	Brain	33-105-3-F5-PU
ID140	new	4	Brain	33-35-2-H11-PU
ID141	new	4	Brain	33-50-3-E12-PU
ID142	new	4	Brain	33-16-3-H7-PU
ID143	new	4	Brain	33-79-2-H4-PU
ID144	new	3.9	Brain	33-32-4-B12-PU
ID145	new	3.9	Brain	33-110-4-A5-PU
ID146	new	3.9	Brain	33-109-2-H1-PU
ID147	new	3.9	Brain	33-100-1-E6-PU
ID148	new	3.9	Brain	33-78-2-E7-PU
ID149	new	3.9	Brain	33-82-4-G3-PU
ID150	new	3.9	Brain	17-1-1-A9-PU
ID151	new	- 3.9	Brain	33-89-4-E1-PU
ID152	new	3.9	Brain	33-89-1-B4-PU
ID153	new	3.9	Brain	33-96-3-A3-PU
ID154	new	3.8	Brain	33-92-3-D1-PU
ID155	new	3.8	Brain	33-104-4-H4-PU
ID156	new	3.8	Brain	33-106-1-B8-PU
ID157	new	3.6	Brain	33-1-3-D1-PU
ID158	new	3.6	Brain	33-40-2-F5-PU
ID159	new	3.6	Brain	33-4-1-E8-PU
D160	new	3.6	Brain	33-36-3-E2-PU
D161	new	3.6	Brain	17-18-3-A6-PU
ID162	new	3.6	Brain	33-12-1-B1-PU
ID163	new	3.6	Brain	33-29-1-H1-PU
ID164	new	3.6	Brain	33-103-1-E1-PU
ID165	new	3.5	Brain	33-10-4-H2-PU
ID166	new	3.5	Brain	33-25-1-H2-PU
ID167	new	3.5	Brain	33-10-4-G2-PU
ID168	new	3.5	Brain	33-67-1-F4-PU
ID169	ext-est-not-vrt	12.5	Brain	33-77-4-E2-PU
ID170	ext-est-not-vrt	10.1	Brain	33-31-3-C11-PU
ID171	ext-est-not-vrt	9.8	Brain	33-28-2-H7-PU
ID172	ext-est-not-vrt	9.2	Brain	33-112-3-C8-PU
ID173	ext-est-not-vrt	7.9	Brain	33-23-3-A11-PU
ID174	ext-est-not-vrt	7.9	Brain	33-29-2-E11-PU
ID175	ext-est-not-vrt	7.9	Brain	33-66-4-C7-PU
ID176	ext-est-not-vrt	7.1	Brain	33-78-1-D7-PU
ID177	ext-est-not-vrt	6.6	Brain	33-31-3-D7-PU
ID178	ext-est-not-vrt	6.3	Brain	33-19-1-C11-PU
ID179	ext-est-not-vrt	6	Brain	33-67-1-A5-PU
ID180	ext-est-not-vrt	5.9	Brain	33-58-3-C8-PU
ID181	ext-est-not-vrt	4.9	Brain	33-107-4-C3-PU
ID182	ext-est-not-vrt	4.9	Brain	33-7-2-G12-PU
ID183	ext-est-not-vrt	4.8	Brain	33-11-1-G5-PU
ID184	ext-est-not-vrt	4.7	Brain	33-31-4-D9-PU
ID185	ext-est-not-vrt	4.6	Brain	33-26-4-E10-PU
ID186	ext-est-not-vrt	4.5	Brain	33-70-4-F7-PU
ID187	ext-est-not-vrt	4.5	Brain	33-19-2-D1-PU
ID188	ext-est-not-vrt	4.4	Brain	33-48-4-F8-PU

SEQ. ID		VON HEIJNE	TISSUE	INTERNAL
<u>NO.</u>	CATEGORY	SCORE	SOURCE	<b>DESIGNATION</b>
ID189	ext-est-not-vrt	4.3	Brain	33-109-3-B10-PU
ID190	ext-est-not-vrt	4.1	Brain	33-30-2-A6-PU
ID191	ext-est-not-vrt	3.8	Brain	33-75-3-D7-PU
ID192	ext-est-not-vrt	3.7	Brain	33-109-4-C1-PU
ID193	est-not-ext	10.5	Brain	33-97-3-D4-PU
ID194	est-not-ext	10.1	Brain	33-61-2-F6-PU
ID195	est-not-ext	9.5	Brain	33-54-1-B9-PU
ID196	est-not-ext	9.3	Brain	33-39-4-D1-PU
ID197	est-not-ext	9.1	Brain	33-57-4-H5-PU
ID198	est-not-ext	9	Brain	33-60-2-B3-PU
ID199	est-not-ext	8.6	Brain	33-52-1-A1-PU
ID200	est-not-ext	8.4	Brain	33-82-2-H10-PU
ID201	est-not-ext	7.5	Brain	33-79-4-B11-PU
ID202	est-not-ext	7.5	Brain	33-18-3-H3-PU
ID203	est-not-ext	7.5	Brain	33-21-1-D6-PU
ID204	est-not-ext	7.4	Brain	33-17-3-F9-PU
ID205	est-not-ext	7.4	Brain	33-70-2-G3-PU
ID206	est-not-ext	7.4	Brain	33-89-3-H4-PU
ID207	est-not-ext	7.4	Brain	33-46-3-E10-PU
ID208	est-not-ext	7	Brain	33-36-2-F9-PU
ID209	est-not-ext	6.8	Brain	33-39-1-C4-PU
ID210	est-not-ext	6.8	Brain	33-65-4-C6-PU
ID211	est-not-ext	6.4	Brain	33-18-2-G6-PU
ID212	est-not-ext	6.4	Brain	33-36-3-C6-PU
ID213	est-not-ext	6	Brain	33-79-2-B6-PU
ID214	est-not-ext	5.9	Brain	33-79-2-80-PU 33-71-4-D11-PU
ID215	est-not-ext	5.9	Brain	17-12-2-A3-PU
ID216	est-not-ext	5.9	Brain	33-95-1-A12-PU
ID217	est-not-ext	5.8	Brain	33-5-3-E3-PU
ID217				· · <del>-</del>
ID218 ID219	est-not-ext	5.8	Brain	33-74-2-D3-PU
ID219 ID220	est-not-ext	5.7	Brain	33-50-3-H8-PU
ID220 ID221	est-not-ext	5.6	Brain	33-19-1-A2-PU
	est-not-ext	5.5	Brain	33-22-1-D3-PU
ID222	est-not-ext	5.5	Brain	33-97-1-G4-PU
ID223	est-not-ext	5.4	Brain	33-65-4-D10-PU
ID224	est-not-ext	5.4	Brain	33-79-4-C4-PU
ID225	est-not-ext	5.3	Brain	33-20-2-C5-PU
ID226	est-not-ext	5.2	Brain	33-34-4-A5-PU
ID227	est-not-ext	5.2	Brain	33-6-2-F11-PU
ID228	est-not-ext	5.2	Brain	33-2-2-G5-PU
ID229	est-not-ext	5.1	Brain	33-98-1-G7-PU
ID230	est-not-ext	5.1	Brain	33-20-3-B10-PU
ID231	est-not-ext	5	Brain	33-106-2-D9-PU
ID232	est-not-ext	4.9	Brain	33-72-2-A9-PU
ID233	est-not-ext	4.9	Brain	33-83-3 <b>-</b> G8-PU
ID234	est-not-ext	4.8	Brain	33-31-3-E6-PU
ID235	est-not-ext	4.7	Brain	33-28-4-E2-PU
ID236	est-not-ext	4.6	Brain	33-101-3-F4-PU
ID237	est-not-ext	4.6	Brain	33-98-4-C1-PU
ID238	est-not-ext	4.5	Brain	33-31-2-E11-PU
ID239	est-not-ext	4.5	Brain	33-26-2-B6-PU

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SEQ. ID		VON HEIINE	TISSUE	INTERNAL
NÔ.	CATEGORY	SCORE	SOURCE	DESIGNATION
<del></del>			GOORCE	DESIGNATION
ID240	est-not-ext	4.4	Brain	33-75-4-H7-PU
ID241	est-not-ext	4.3	Brain	33-13-1-C6-PU
ID242	est-not-ext	4.3	Brain	33-35-4-G1-PU
ID243	est-not-ext	4.3	Brain	33-76-3-G11-PU
ID244	est-not-ext	4.2	Brain	33-72-1-A3-PU
ID245	est-not-ext	4.2	Brain	33-71-2-A2-PU
ID246	est-not-ext	4.2	Brain	33-23-3-H10-PU
ID247	est-not-ext	4.2	Brain	33-13-1-C1-PU
ID248	est-not-ext	4.2	Brain	33-43-2-G12-PU
ID249	est-not-ext	4.2	Brain	33-91-4-E10-PU
ID250	est-not-ext	4.1	Brain	33-113-2-B8-PU
ID251	est-not-ext	4	Brain	33-104-3-G9-PU
ID252	est-not-ext	3.9	Brain	33-66-2-B10-PU
ID253	est-not-ext	3.9	Brain	33-1-2-E9-PU
ID254	est-not-ext	3.9	Brain	33-51-1-G7-PU
ID255	est-not-ext	3.9	Brain	33-32-3-D11-PU
ID256	est-not-ext	3.8	Brain	33-43-2-H10-PU
ID257	est-not-ext	3.8	Brain	33-48-4-H11-PU
ID258	est-not-ext	3.8	Brain	33-8-4-C5-PU
ID259	est-not-ext	3.8	Brain	33-24-1-F5-PU
ID260	est-not-ext	3.8	Brain	33-70-1-A9-PU
ID261	est-not-ext	3.8	Brain	33-30-4-C4-PU
ID262	est-not-ext	3.8	Brain	33-10-2-G7-PU
ID263	est-not-ext	3.6	Brain	33-18-4-E12-PU
ID264	est-not-ext	3.6	Brain	33-52-1-G7-PU
ID265	est-not-ext	3.6	Brain	33-57-1-H10-PU
ID266	est-not-ext	3.5	Brain	33-80-3-E2-PU
ID267	est-not-ext	3.5	Brain	33-36-1-D3-PU
ID268	ext-vrt-not-genomic	11.3	Brain	33-101-1-A2-PU
ID269	ext-vrt-not-genomic	6.6	Brain	33-55-2-E8-PU
ID270	ext-vrt-not-genomic	4.8	Brain	33-14-2-H3-PU
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### TABLE III

SEQ. ID	
NO.	SIGNAL PEPTIDE
ID38	MLLLLGLCLGLSLC
ID39	MENGGAGTLQIRQVLLFFVLLGMSQA
ID40	MRGPEPGPQPTMEGDVLDTLEALGYKGPLLEEQALTKAAEGGLSSPEFSELCIWLGSQIK
	SLCNLEESITSAGRDDLESFQLEISGFLKEMACPYSVLISGDIKDRLKKKEDCLKLLLFL
	STELQA
ID41	MEKSWMLWNFVERWLIALASWSWALC
ID42	MQQTRTEAVAGAFSHCLGFCGMRLGLLLLARHWCIA
ID43	MEKGNAFLKNRLVVFLLLPLASGP
ID44	MFPFNQAGLPTLLMLIVFHAASMA
ID45	MTSRSLRRCSCLRVTHNKEILASTVSLGVEGYMLGGGSRINSSNLNDGEEECSPDSLLVW
ID46	KKKSLLLWMSSLPSLG
ID46	MWTASAMDFRTCIASXLPALCYVQACRALMIAASVLGLPAILLLLTVLPCIXM
ID47 ID48	MGPPPTHIKYLHLNIYCNGKSTAPGIRSHSLGFALLSLSHPTCQA
ID48 ID49	MFCLLTFLAFTTLLFA
ID50	MHCGSTPGLCPCWVPFLKCLLAVLSSLFA
ID50 ID51	MNLVCSALLLLGIVSS MSVI DDBODDI VIVOKDUSSI EAANG IGU AUG OFF
ID51	MSVLDDRQRDILVVQKRHSSLEAAMLIGLLAWLQT MGVNGRRLLIICHYLPLSLC
ID53	· · · · · · · · · · · · · · · · · · ·
ID54	MKLRECPALRWSQLSQHKLECLLLYLAESSG MDPRGILKAFPKRQKIHADASSKVLAKIPRREEGEEAEEWLSSLRAHVVRTGIGRARAEL
1034	FEKQIVQHGGQLCPAQGPGVTHIVVDEGMDYERALRLLRLPQLPPXCSA
ID55	MFWKLSLSLFLVAVLVKVAEA
ID56	MAFLGLFSLLVLQSMATG
ID57	MAFLGLFSLLVLQSMATG
ID58	MSFSLNFTLPANTTSSPVTGGKETDCGPSLGLAAGIPLLVATALLVALLFTLIHR
ID59	MSTWYLALNKSYKNKDSVRIYLSLCTVSIKFTYFHDIQTNCLTTWKHSRCRFYWAFGGSI
	LQHSVDPLVLFLSLALLVTP
ID60	MAIGISLQLLCCIFTLVLQ
ID61	MQATSNLLNLLLLSLFAGL
ID62	MMKWKPEDLGSVPCEAFSVTLLCGWPGSHWC
ID63	MQATSNLLNLLLLSLFAGL
ID64	MASSHWNETTTSVYQYLGFQVQKIYPFHDNWNTACFVILLLFIFTVVS
ID65	MLWFSGVGALAERYCRRSPGITCCVLLLLNCSG
ID66	MLFLQMGKQSWTLIFFLNVTQLVRG
ID67	MELRXXPPGGREVQLLLGLCSPPXXSL
ID68	MLWSLLSSSGSHFG
ID69	MDISGLIPGLVSTFILLSXSDHYGRKFPMILSSVGALATSVWLCLLCYFAFP
ID70	MXVFFSKNRFEMYFSLLLFVILLITSLIFC
ID71	MPVPACWISSSLSLLASHHSVSC
ID72	MCPVFSKQLLACGSLLPGLWQ
ID73	MALTIHGERMRPDWESPWITSSQAQSLSLGGSPSSRGPLVPRGEYLASCPEGVRSHSHLL
	PRSLLPLSAWPPWAWH
ID74	MAARFRCGHLCVPEVPRGPASHAEGGGGRLSRKAAHQAQLCWRAGGDGRGNFN
	PMNFLVAGTFASSCHSPPLLWSLPPRILIASSLPTLSHP
ID75	MASTISAYKEKMKELSVLSLICSCFYTQP
ID76	MLQVYGKPVYQGHRSTLKKGPYLRFNSPSPKSRPQRPKVIERVKGTKVKSIRTQTDFYAT
	KPKKMDSKMKHSVPVLPHGDQQYLFSPSREMPTFSGTLEGHLIPMAILLGQTQS
ID77	MSVLEISGMIMNRVNSHIPGIGYQIFGNAVSLILGLTPFVFRLSQATDLEQLTAHSASEL
	YVIAFGSNEDVIVLSMVIISFVVRVSLVWIFFFLLCVAERTYKQRLLFAKLFGHLTSA

SEQ. ID	
<u>NO.</u>	SIGNAL PEPTIDE
W-50	1
ID78	MCKGIKAGDTCEKLVGYSAVYRVCFGMACFFFIFCLLTLKINNSKSCRAHIHNGFWFFKL
man	LLLGAMCSG
ID79	MSDSAGGRAGLRRYPKLPVWVVEDHQEVLPFIYRAIGSKHLPASNVSFLHFDSHPDLLIP
ID80	VNMPADTVFDKETLFGELSIENWIMPAVYAGHFSHVVWFHPTWA
ID81	MSSCRGQKVAGGLRVVSPFPLCQPAGEPSRGKMRSSCVLLTALVALA
ID81 ID82	MIIPFKIKNLGGRVLLSGREMFPASVRAPDLAVALSLLPAWT
ID82	MVCSAPRKIVVRAFITIIFIYYAIKKRANEPAAYLMLKPEALILLLLAQKGPS
ID84	MTESSMKKLASTLLDAITDKDPLVQEQVCSALCSLGEVRP
ID85	MQETDCNKRWGRGLGGLWSETGRRFHCKSFVFLFHCTSGLSSC MLLEVPWLSSTVSCAQG
ID86	MSGGRMQARCSQQSTWSPAFLAVAGPGWA
ID87	MLQMLWHFLASFFPRAGC
ID88	MYSHPVSSLVCLLAMGKGLG
ID89	MGRKEEDDCSXWKKQTTNIRKTFIFMEVLGSGAFS
ID90	MMIAVFGNANDRNVLTLLPNQSLFSLARA
ID91	MFFELPLVVTAWFFGMCRS
ID92	MNHNIICVMYIVPFLMTKCLYFCHSCKRGSFLLIVANVHFSQT
ID93	MSCGSAASLTGLCXCCLQALG
ID94	MQAVDNLTSAPGNTSLCTRDYKITQVLFPLLYTVLFFVGLITNGLA
ID95	MAAAMXLLCSSCCSWGPAAG
ID96	MDFIKDQSLSHRSVVKVLSLRKAQA
ID97	MTRPFWASCSTWATSRISCAFSLASSTA
ID98	MKSCAVSLTTAAVAFG
ID99	MSIHECACLSLSLICLRMSLS
ID100	MLSGLSFLSVFSLWC
ID101	MGLKDKSQAPASGLGVLRGQRSGSFISMPAPASGQXPEESRSPAPPVASRSQNRGYRPWH
	GPLWVHQSVRFGLYSILHFPFWVHG
ID102	MSDQIKFIMDSLNKEPFRKNYNLITFDSLEPMQLLQVLSDVLA
ID103	MSPSCLHPDLWSMCLEVPSFTATDSVNCGCCLELATEPARNIRSTTRASLLRCSSFTSTR
	NSTGISALPPAAPMAWPFSASLSTLPVPLTHSSVASLTATPSLA
ID104	MDLSFHLLLDPSSTQS
ID105	MPHFLDWFVXVYLVISVLILVGFGAC
ID106	MSKLKVIPEKSLTNNSRIVGLLAQLEKINA
D107	MMSASRLAGTLIPAMAFLSCVRP
ID108	MVDGTQLRGLTRMYQVPLXLDRDETLVRLRFTMVALVTVCCXLVAFLFC
ID109	MKQNFLVLNSVWYLISMLQMLAVIIT
ID110	MECQNSSLKKCLLVEKSLVKASYLIAFQTAASKKPFSIAEELIKPYLVEMCLEVLGSSA
D111	MHSSIKTKGSVMWLVALLEMCVC
ID112 ID113	MTVLPLEAISSLSSFVLG
	MGTASRSNIARHLQTNLILFCVGAVGACTL
ID114 ID115	MNSSKEEMRELAALFYSVVVSTVSG
D115	MSQDGGXGELKHMVMSFRVSELQVLLGFAGRNKSGRKHELLAKALHLLKSSC
ID117	MPCISLLGLLYNFVQVLCYLSIFCLGVLF MKIAVLFCFFLLIIF
ID118	
ID118	MAKQKPHVLGSRVMPASCVSERRRKPSFQVSTWSSASLRGSWQ MGFLYLKSVFDVSLG
ID120	MRMGPGRKRDFSPVPWSQYFESMEDVEVENETGKDTFRVYKSGSEGPVLLLLHGGG
	HSALS
ID121	MIFLLYLLPSSEE
ID122	MRMGPGRKRDFSPVPWSQYFESMEDVEVENETGKDTFRVYKSGSEGPVLLLLHGGG
	HSALS

SEQ. ID	
<u>NO.</u>	SIGNAL PEPTIDE
ID123	MLSLLNLISILASIPS
ID124	MGTTSNMVTTIHLMLLWPVHPLLVG
ID125	MGDPERPEAAGLDQDERSSSDTNESEIKSNEEPLLRKSSRRFVIFPIQYPDIWKMYKQAQ
	ASFWTAEEVDLSKDLPHWNKLKADEKYFISHILAFFAASDG
ID126	MDAGLFSLLPHPPCVG .
ID127	MLITLTYLIQGESA
ID128	MYTGFRIEATLLTRVQCLCAIPFAFS
ID129	MYKQAQASFWTAEEVDLSKDLPHWNKLKADEKYFISHILAFFAASDG
ID130	MLLHLCSVKNLYQNRFLGLAAMASPSRN
ID131	MPCPTWTCLKSFPSPTSS
ID132	MEDLFSPSIXPPAPNISVPILLGWGLNLTLGQG
ID133	MAETKDAAQMLVTFKDVAVTFTREEWRQLDLAQRTLYREVMLETCGLLVSLG
ID134	MLILSQNIAQLEA
ID135	MLLGASAQGLWAHSWTCSCSA
ID136	MAAPLELSCWGGGWG
ID137	MSXVGIDLGFLNCYIAVARS
ID138	MEYSKXFVVFSTMFTASSP
ID139	MPMASSPPPSPHPQEPAPLLPSLPRLSLPFRLPWASTATA
ID140	MQHVXGHXPDPIAIMYVCPPCGHTTWALGLKFLSSSSQ
ID141	MGWEMTCIKSFFWARSHAGFLKCLLLSSLQ
ID142	MVFGGVCPSVTSIIAESLQGWNLVQLSFAATTPVLA
ID143	MHFITWSLLFLYQCSL
ID144	MSGASPIERTPMEEAPSSCPTSSCWPSVASPSSSWS
ID145	MEWAGKQRDFQVRAAPGWDHLASFPGPSLRLFSGSQA
ID146	MIAFFDEDNPRKRRSYSFTQSAGILCQETTYSTPHTKLEKAKSPTADAKVVSLSLQTSSA
ID147	MGKSIXSLCSVXLKARLKGXLEAVHLCLRAQKRRTALFCTLPCPVERG
ID148	MCLHMTLFRVPFTFS
ID149	MLNILKTLTSAALP
ID150	MRARVWPRSHGIPVPSFLSKSSLSHTPSPLLCLYHPPVYT
ID151	MWNAVAIICNGSWCQTXSTSGLESLCLSLLIPGPKP
ID152	MLRLGLFKISWARC
ID153	MPFAEDKTYKYICRNFSNFCNVDVVEILPYLPCLTA
ID154	MPGSSGLRFICKSRNHPQFGSFSGTDSLSFLPPCPC
ID155	MDVTGDEEEEIKQEINMLKKYSHHRNIATYYGAFIKKNPPGMDDQLXLVMEFCGAGS
ID156	MIFGLYFVLAVKLFLVFLLNICKG
ID157	MRKKRVEELIVFPGEVTSFSSIKCSSWISSLASG
ID158	MPSSSLAELCLMQQDACLFSXFLAVSRH
ID159	MDLWSCLFPVMLMEPSKGLEDSEWKMALQMRMQLPCLVLG
ID160	MSGKGKCRPIALRRAVPLPTTSTLTSA
ID161	MTPKAIQKSSGLFCPSQA
ID162	MPDQFDQAVVLNQLRYSGMLETVRIRKAGYAVRRPFQDFYKRYKVLMRNLALPEDV
	RGKCTSLLQLYDASNS
ID163	MCLVSFFLELNVLQQ
ID164	MRSLACLTPCGHA
ID165	MHLLSNWANPASS
ID166	MWSGKWALVSPFAMLHSVWRLIPA
ID167	MKVHMHTKFCLICLLTFIFH
ID 168	MGRRHWVLTHSALSLFYTADTSHG
ID169	MAVFVVLLALVAGVLG
ID170	MAPLLLQLAVLGAALA

SEQ. ID	
NÔ.	SIĞNAL PEPTIDE
ID171	MPVTVTRTTITTTTSSSGLGSPMIVGSPRALTQPLGLLRLLQLVSTCVA
ID172	MELVLVFLCSLLAPMVLA
ID173	MGPIWSSYYGNCRSLLFVMDASDPTQLSASCVQLLGLLSAEQLAEA
ID174	MSGGRAPAVLLGGVASLLLSFVWMPALLPVASRLLLLPRVLLTMASG
ID175	MALSCTLNRYLLLMAQEHLEFRLPEIXSLLLLFGGQFASS
ID176	MAARGVIAPVGESLRYAEYLQPSAKRPDADVDQQRLVRSLIAVGLGVAALAFA
ID177	MRMCAGSIYKSATQAVLGXLFLGGLCRG
ID178	MAERRPLSPIPSXRRPSEPSRPRPAAAGXRSLPRPGDEELQLPCAVHDLIFWRDVKKTG
TD170	FVFGTTLIMLLSWQLSVS
ID179	MAAPVLLRVSVPRWERVARYAVCAAGILLSIYAYHVEREKERDPEHRALCDLGPWVK
ID180	CSAALASRWGRGFGLLGSIFGKDGVLNQPNSVFGLIFYILQLLLGMTASAVA
ID181	MSFLQDPSFFTMGMWSIGAGALGAAALALLLANT
ID182	MASLLCCGPKLAACGIVLSAWGVIMLIMLGIFFNVHS MILPYRMXSLFLHAVSSSFT
ID183	
Шібэ	MATLVELPDSVLLEIFSYLPVRDRIRISRVCHRWKRLVDDRWLWRHVDLTLYTVRALAGR AWA
ID184	MKNACIVLPPTPPPSLQPSASLLAPNRFLFSCFCFLSHKFG
ID185	MAFGLQMFIQRKFPYPLQWSLLVAVVAG
ID186	MYCKILVLMLHTELIRTDYSSVDQLLLNYPAEEGLGRERSLLWTPLLSPGSLR
ID187	MAVSHSVKERTISENSLIILLQGLQG
ID188	MESGGRPSLCQFILLGTTSVVTA
ID189	MAALDLRAXWIRWSCSCLGXLXGAGGETNGVERPGGGGLALARQGSLRDGRQVGR
	APAVCFPHGAPGLPPRQRXXGGXPEVQGGESWCPRPRGGGASRTGLRRRKGPTKTPE
	PESSEAPQDPLNWFGILVPHSLRQAQA
ID190	MAFLPSPAWWISLLPSLLSIC
ID191	MEPKVAELKQKIEDTLCPFGFEVYPFQVAWYNELLPPAFHLPLPGPTLA
ID192	MLVLRSGLTKALA
ID193	MSGGHLADLTLLFVLLLFSLLPA
ID194	MKPSRTPARLWMLPQQQAGAVVVAAPTERHPTHHMAGWLLGALTLLGLVTS
ID195	MGESIPLAAPVPVEQAVLETFFSHLGIFSYDKAKDNVEKEREANKSAGGSWLSLLAALAH
	LAAA
ID196	MQMSYAIRCAFYQLLLAALMLVAMLQLLYLSLLSGLHG
ID197	MLRAELKIAVVLFAFHLLLSFILG
ID198	MNHQQTLIGRLLCDLHGLSLSPPVANNVQALFRMLTPEAYSCLLILLLRTFLCSA
ID199	MIITAVVSISVTIFCFQTKVDFTSCTGLFCVLGIVLLVTG
ID200	MAAGGRMEDGSLDITQSIEDDPLLDAQLLPHHSLQAHFRPRFHPLPTVIIVNLLWFIHLV
ID201	FVVLX MCDCOM LEMECENCO
ID201 ID202	MSPGCMLLFVFGFVGG MKLLLGIALLAYVAS
ID202	MDILVPLLQLLVLLLTLPLHLMA
ID204	MEAASPSNSTGVERXADLMDADSLLLSLELASGSG
ID205	MIRQERSTSYQEAVRPALPSSKPCLLTSPAVLVKLLSSSASTS
ID206	MKLIDYGLSGYQEESAEVKAMDFITSTAILPLLFGCLGVFG
ID207	MRCLTTPMLLRALAQAARA
ID208	MSRFLNVLRSWLVMVSIIAMGNTLQSFRDHTFLYEKLYTGKPNLVNGLQARTFGIWTLLS
- <del></del> - <del>-</del> <del>-</del> -	SVIRCLC
ID209	MIFLTLSLDSRVSA
ID210	MQCFSFIKTMMILFNLLIFLCGAALLXVG
ID211	MAEAALEAVRXSYENSRPLQGSSACLLLCPTWTNP
ID212	MATASPSVFLLMVNGQVES
ID213	MAGIKALISLSFGGAIGLMFLMLGCALP

SEQ. ID	·
NO.	SIGNAL PEPTIDE
ID214	MIGDILLFGTLLMNAGA
ID215	MKTMILTLSLFGSCIS
ID216	MDWRVPPSXXDPGHQDIPLPVTXXFISVSVLSSLGIVLA
ID217	MAAAALPAWLSLQSRA
D218	MAMVSAMSWVLYLWISACAMLLCHG
D219	MGKEWGWQEMENGGAAPAWGAGPPYHPAPPPVEKTLSWGCGFGLHSGFGGSGGG
	VGLCRLLCLVRLFCC
ID220	MLQTSNYSLVLSLQFLLLSYD
D221	MWFEILPGLSVMGVCLLIPGLATA
D222	MRPSPLSGILADPLXLFPFSEG
D223	MRESLSXRSWHLPASLMMAQXFIPAVA
D224	MSGVVPTAPEQPAXEMENQTKPPDPRPDAPPEYSSHXFTRTPWKQLSLHLLATRACYG
D225	MWRYQFGWGVITRGPREIPFPPSLLASESLLPPLPDLVLTCTSLGFVTRVWMSLNLNELS
	LYSRTWVFTCLVFFCFG
D226	MVKLLVAKILCMVGVFFFMLLGSLLPVKI
D227	MPVSIMCLIGLKANASS
D228	MKVILLYLVLEKLVSRA
D229	MAVTLSLLLGGRVCXPSLA
D230	MLNQTSGRTSLLPELGVVTPAQG
D231	MTSENLVQTAPKKKKNKGKKGLEPSQSTAAKVPKKAKTWIPEVHDQKADVSAWKDL
	FVPRPVLRALSFLGFSAPTPIQA
D232	MAAFGRQXXXWHXLIPLTWACMA
D233	MSLTSSPKKRRSICFDRFLMPQSQSGPSSLGESYRTGVGFLIPEGWFLSGCPHGSSA
D234	MGELGNRSRCILFLSENPCLSESIFQSLXFCLSPPPSPS
D235	MAELGLNEHHQNEVINYMRFARSKRGLRLKTVDSCFQDLKESRLVEDTFTIDEVSEVLNG
	LQAVVHSEVESELINTAYTNVLLLRQXFAQAEK
D236	MVTLPSGTWAFSCPYLALVDGGMLGSAREDAHASVVSWAVGLLYAVAQG
D237	MASASARGNQDKDAHFPPPSKQSLLFCPKXXLHIHRAEISKIMRECQEESFWKRALPFSL VSMLVTQG
D238	MLLMKSILLKVVCVLCIYLKFKLMALIYVPDKNNTNNNILRYNHNEISIGISVQCHFILS
	LCVLCIVLT
D239	MAQRLLLRRFLASVIS
D240	MAASKVKQDMPPXGGYGPIDYKRNLPRRGLSGYSMLAIGIGTLIYGHWSIMKWNRERRRL
	QIEDFEARIALLPLLOA
D241	MRHLVTEELFPCSNLEDVVEDNSHSYFTLRITMACKGVPSTLLSLAILSHISTP
D242	MSAEVKVTGQNQEQFLLLAKSAKGAALATLIHQVLEAPGVYVFGELLDMPNVRELAESXF
	ASTFRLLXVFAYGTYA
D243	MLLSIGMLMLSATQVXTILXVQLFAFLNLLPVEA
D244	MGWEVVSLSYCGVSWG
D245	MRECISVHVGQAGVQIGNACWELFCLEHGIQA
D246	MAGPLQGGGARALDLLRGLPRVSLA
D247	MPAGVPMSTYLKMFAASXLAMCAGA
D248	MAVQCVRLARRSLPALALSLRASP
D249	MFSIISRSRACSMYFKENAKPSQLRLMHHYLSTPTSA
D250	MKRLLPATSLAGPVLS
D251	MLIITNPWPKYFDAAGRLTPEFSQRLTNKIRELLQQMERGLKSADXXDGTGYTGWAGIAV
	LYLHLYDVFG
D252	MCATETVRAWLAQGSSSAGWG
D253	MLLLATHPETVGQVTLRVXPVSLEVSIQMCAAAAAAFCLKXXGANT
ID254	MAASSATPAPXXSQRCGADAGSAARIVFRWGRGRRGARSPEGSGHHGRANSGLGGAQ
	LOGGAXG

SEQ. ID _NO	SIGNAL PEPTIDE
ID255	MLRRPLAGLAAAALGRA
ID256	MDRPGFVAALVAGGVAG
ID257	MIVWFEGISMDLLTLLFQRRS
ID258	MRTFVHFALDALMFPARRRA
ID259	MAAPPQLRALLVVVNALLRKRRYHAALAVLKGFRNGAVYGAKIRAPHALVMTFLFR
	NGSLQ
ID260	MPVDLGXALGLLPSLAKA
ID261	MNLFIMYMAGNTISIFPTMMVCMMAWRPIQALMAISATFKMLESSSQKFLQGLVYLIGNL
	MGLALAVYKCQS
ID262	MISLTDTQKIGMGLTGFGVFFLFFGMILFFDKALLAIGNVLFVAGLAFVIG
ID263	MAASGAPRILVDLLKLXVAPLAVFQMLKSMCAG
ID264	MASVSSATFSGHGARSLLQFLRLVGQ
ID265	MWYLAVLLVLFTLNIL
ID266	MFTFGRLFQIITVVTCLQFIQDCCIHSRQINSLLEXSSLSRC
ID267	MIQDRDRCAQAAAVAAVGNLEPRGTPGPEDEAFCLPGCVGTLCQLDWWIWG
ID268	MKIIFPILSNPVFRRTVKLLLCLLWIGYSQG
ID269	MVSRMVSTMLSGLLFWLASGWTPAFA
ID270	MTATLAAAADIATMVSGSSGLAXA

Minimum signal peptide score	false positive rate	false negative rate	proba(0.1)	proba(0.2)
3.5	0.121	0.036	0.467	0,664
4	0.096	0.06	0.519	0.708
4.5	0.078	0.079	0.565	0.745
5	0.062	0.098	0.615	0.782
5.5	0.05	0.127	0.659	0.813
6	0.04	0.163	0.694	0.836
6.5	0.033	0.202	0.725	0.855
7	0.025	0.248	0.763	0.878
7.5	0.021	0.304	0.78	0.889
<b>]</b> 8	0.015	0.368	0.816	0.909
8.5	0.012	0.418	0.836	0.92
9	0.009	0.512	0.856	0.93
9.5	0.007	0.581	0.863	0.934
10	0.006	0.679	0.835	0.919

**TABLE IV** 

Minimum signal peptide score		New ESTs	ESTs matching public EST closer than 40 bp from beginning	ESTs extending known mRNA more than 40 bp	ESTs extending public EST more than 40 bp
3.5	2674	947	599	23	150
4	2278	784	499	23	126
4.5	1943	647	425	22	112
5	1657	523	353	21	96
5.5	1417	419	307	19	80
6	1190	340	238	18	68
6.5	1035	280	186	18	60
7	893	219	161	15	48
7.5	753	173	132	12	
8	636	133	101	11	29
8.5	543	104	83	1	26
9	456		63		24
9.5	364	57	48	_	18
10	303	47	35	6	15

**TABLE V** 

			ESTs	ESTs	ESTs
			matching	extending	extending
Tissue	All ESTs	New ESTs	public EST	known	•
113300		148M E212	closer than		public EST more than 40
			40 bp from		•
			beginning	than 40 bp	bp
Brain	329	131	75	3	24
Cancerous prostate	134	40	37	1	6
Cerebellum	17	9	1	0	6
Colon	21	11	4	0	o
Dystrophic muscle	41	18	8	0	1
Fetal brain	70	37	16	0	1
Fetal kidney	227	116	46	1	19
Fetal liver	13	7	2	0	o
Heart	30	15	7	0	11
Hypertrophic prostate	86	23	22	2	2
Kidney	10	7	3	0	o
Large intestine	21	8	4	0	1
Liver	23	9	6	0	ol
Lung	24	12	4	0	1
Lung (cells)	57	38	6	0	4
Lymph ganglia	163	60	23	2	12
Lymphocytes	23	6	4	0	2
Muscle	33	16	6	0	4
Normal prostate	181	61	45	7	11
Ovary	90	57	12	1	2
Pancreas	48	11	6	0	1
Placenta	24	5	1	0	4
Prostate	34	16	4	Ō	
Spleen	56	28	10	0	
Substantia nigra	108	47	27	1	6
Surrenals	15	3	3	1	Ō
Testis	131	68	25	1	8
Thyroid	17	8	2	Ö	2
Umbilical cord	55	17	12	-	
Uterus	28	15	3	0	
Non tissue-specific	568	48	177	-	
Total	2677	947	601	23	

**TABLE VI** 

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## Description of Transcription Factor Binding Sites present on promoters isolated from SignalTag sequences

Promoter sequence P13H2 (646 bp):

Matrix	Position	Orientation	Score	Length	Sequence
CMYB_01	-502	•	0.983	9	TGTCAGTTG
MYOD_Q6	-501	•	0.961	10	CCCAACTGAC
S8_01	-444	•	0.960	11	AATAGAATTAG
S8_01	-425	•	0.966	11	AACTAAATTAG
DELTAEF1_01	-390	•	0.960	11	GCACACCTCAG
GATA_C	-364	•	0.964	11	AGATAAATCCA
CMYB_01	-349	•	0.958	9	CTTCAGTTG
GATA1_02	-343	•	0.959	14	TTGTAGATAGGACA
GATA_C	-339	•	0.953	11	AGATAGGACAT
TAL1ALPHAE47_01	-235	+	0.973	16	CATAACAGATGGTAAG
TAL1BETAE47_01	-235	+	0.983	16	CATAACAGATGGTAAG
TAL1BETAITF2_01	-235	•	0.978	16	CATAACAGATGGTAAG
MYOD_Q6	-232	•	0.954	10	ACCATCTGTT
GATA1_04	-217	•	0.953	13	TCAAGATAAAGTA
IK1_01	-126	+	0.963	13	AGTTGGGAATTCC
IK2_01	-126	+	0.985	12	AGTTGGGAATTC
CREL_01	-123	+	0.962	10	TGGGAATTCC
GATA1_02	-96	+	0.950	14	TCAGTGATATGGCA
SRY_02	-41	•	0.951	12	TAAAACAAAACA
E2F_02	-33	+	0.957	8	TTTAGCGC
MZF1_01	-5	•	0.975	8	TGAGGGGA

### Promoter sequence P16B4 (861bp):

Matrix	Position	Orientation	Score	Length	Sequence
NFY_Q6	-748	•	0.956	11	GGACCAATCAT
MZF1_01	-738	+	0.962	8	CCTGGGGA
CMYB_01	-684	+	0.994	9	TGACCGTTG
VMYB_02	-682	•	0.985	9	TCCAACGGT
STAT_01	-673	•	0.968	9	TTCCTGGAA
STAT_01	-673	•	0.951	9	TTCCAGGAA
MZF1_01	-556	•	0.956	8	TTGGGGGA
IK2_01	-451	•	0.965	12	GAATGGGATTTC
MZF1_01	-424	+	0.986	8	AGAGGGGA
SRY_02	-398	•	0.955	12	GAAAACAAAACA
MZF1_01	-216	•	0.960	8	GAAGGGGA
MYOD_Q6	-190	•	0.981	10	AGCATCTGCC
DELTAEF1_01	-176	+	0.958	11	TCCCACCTTCC
S8_01	5	•	0.992	11	GAGGCAATTAT
MZF1_01	16	•	0.986	8	AGAGGGGA

### Promoter sequence P29B6 (555 bp):

Matrix	Position	Orientation	Score	Length	Sequence
ARNT_01	-311	•	0.964	16	GGACTCACGTGCTGCT
NMYC_01	-309	•	0.965	12	ACTCACGTGCTG
USF_01	-309	+	0.985	12	ACTCACGTGCTG
USF_01	-309	•	0.985	12	CAGCACGTGAGT
NMYC_01	-309	-	0.956	12	CAGCACGTGAGT
MYCMAX_02	-309	-	0.972	12	CAGCACGTGAGT
USF_C	-307	+	0.997	8	TCACGTGC
USF_C	-307	•	0.991	8	GCACGTGA
MZF1_01	-292	-	0.968	8	CATGGGGA
ELK1_02	-105	+	0.963	14	CTCTCCGGAAGCCT
CETS1P54_01	-102	+	0.974	10	TCCGGAAGCC
AP1_Q4	-42	•	0.963	11	AGTGACTGAAC
AP1FJ_Q2	-42	•	0.961	11	AGTGACTGAAC
PADS_C	45	+	1.000	9	TGTGGTCTC

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#### <u>CLAIMS</u>

- 1. A purified or isolated nucleic acid comprising the sequence of one of SEQ ID NOs: 38-270 or comprising a sequence complementary thereto.
  - 2. The nucleic acid of Claim 1, wherein said nucleic acid is recombinant.
- 3. A purified or isolated nucleic acid comprising at least 10 consecutive bases of the sequence of one of SEQ ID NOs: 38-270 or one of the sequences complementary thereto.
- 4. A purified or isolated nucleic acid comprising at least 15 consecutive bases of one of the sequences of SEQ ID NOs: 38-270 or one of the sequences complementary thereto.
  - 5. The nucleic acid of Claim 4, wherein said nucleic acid is recombinant.
  - 6. A purified or isolated nucleic acid of at least 15 bases capable of hybridizing under stringent conditions to the sequence of one of SEQ ID NOs: 38-270 or one of the sequences complementary to the sequences of SEQ ID NOs: 38-270.
    - 7. The nucleic acid of Claim 6, wherein said nucleic acid is recombinant.
  - 8. A purified or isolated nucleic acid encoding a human gene product, said human gene product having a sequence partially encoded by one of the sequences of SEQ ID NO: 38-270.
  - 9. A purified or isolated nucleic acid having the sequence of one of SEQ ID NOs: 38-270 or having a sequence complementary thereto.
    - A purified or isolated nucleic acid comprising the nucleotides of one of SEQ
       NOs: 38-270 which encode a signal peptide.
  - 11. A purified or isolated polypeptides comprising a signal peptide encoded by one of the sequences of SEQ ID NOs: 38-270.
    - 12. A vector encoding a fusion protein comprising a polypeptide and a signal peptide, said vector comprising a first nucleic acid encoding a signal peptide encoded by one of the sequences of SEQ ID NOs: 38-270 operably linked to a second nucleic acid encoding a polypeptide.
- 30 13. A method of directing the extracellular secretion of a polypeptide or the insertion of a polypetide into the membrane comprising the steps of:

obtaining a vector according to Claim 12; and

introducing said vector into a host cell such that said fusion protein is secreted into the extracellular environment of said host cell or inserted into the membrane of said host cell.

- 14. A method of importing a polypeptide into a cell comprising contacting said cell with a fusion protein comprising a signal peptide encoded by one of the sequences of SEQ ID NOs: 38-270 operably linked to said polypeptide.
- 15. A method of making a cDNA encoding a human secretory protein that is partially encoded by one of SEQ ID NOs 38-270, comprising the steps of:

obtaining a cDNA comprising one of the sequences of SEQ ID NOs: 38-270;

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contacting said cDNA with a detectable probe comprising at least 15 consecutive nucleotides of said sequence of SEQ ID NO: 38-270 or a sequence complementary thereto under conditions which permit said probe to hybridize to said cDNA;

identifying a cDNA which hybridizes to said detectable probe; and isolating said cDNA which hybridizes to said probe.

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- 16. An isolated or purified cDNA encoding a human secretory protein, said human secretory protein comprising the protein encoded by one of SEQ ID NOs 38-270 or a fragment thereof of at least 10 amino acids, said cDNA being obtainable by the method of Claim 15.
- 17. The cDNA of Claim 16 wherein said cDNA comprises the full protein coding sequence partially included in one of the sequences of SEQ ID NOs: 38-270.
- 18. A method of making a cDNA comprising one of the sequences of SEQ ID NOs: 38-270, comprising the steps of:

contacting a collection of mRNA molecules from human cells with a first primer capable of hybridizing to the polyA tail of said mRNA:

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hybridizing said first primer to said polyA tail;

reverse transcribing said mRNA to make a first cDNA strand;

making a second cDNA strand complementary to said first cDNA strand using at least one primer comprising at least 15 nucleotides of one of the sequences of SEQ ID NOs 38-270; and

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isolating the resulting cDNA comprising said first cDNA strand and said second cDNA strand.

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- 19: An isolated or purified cDNA encoding a human secretory protein, said human secretory protein comprising the protein encoded by one of SEQ ID NOs 38-270 or a fragment thereof of at least 10 amino acids, said cDNA being obtainable by the method of Claim 18.
- 20. The cDNA of Claim 19 wherein said cDNA comprises the full protein coding sequence partially included in one of the sequences of SEQ ID NOs: 38-270.
- 21. The method of Claim 18, wherein the second cDNA strand is made by: contacting said first cDNA strand with a first pair of primers, said first pair of primers comprising a second primer comprising at least 15 consecutive nucleotides of one of the sequences of SEQ ID NOs 38-270 and a third primer having a sequence therein which is

performing a first polymerase chain reaction with said first pair of nested primers to generate a first PCR product;

included within the sequence of said first primer;

contacting said first PCR product with a second pair of primers, said second pair of primers comprising a fourth primer, said fourth primer comprising at least 15 consecutive nucleotides of said sequence of one of SEQ ID NO:s 38-270, and a fifth primer, said fourth and fifth primers being capable of hybridizing to sequences within said first PCR product; and

performing a second polymerase chain reaction, thereby generating a second PCR product.

- 22. An isolated or purified cDNA encoding a human secretory protein, said human secretory protein comprising the protein encoded by one of SEQ ID NOs 38-270, or a fragment thereof of at least 10 amino acids, said cDNA being obtainable by the method of Claim 21.
- 23. The cDNA of Claim 22 wherein said cDNA comprises the full protein coding sequence partially included in one of the sequences of SEQ ID NOs: 38-270.
- 24. The method of Claim 18 wherein the second cDNA strand is made by: contacting said first cDNA strand with a second primer comprising at least 15 consecutive nucleotides of the sequences of SEQ ID NOs: 38-270;

hybridizing said second primer to said first strand cDNA; and extending said hybridized second primer to generate said second cDNA strand.

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- 25. An isolated or purified cDNA encoding a human secretory protein, said human secretory protein comprising the protein partially encoded by one of SEQ ID NOs 38-270 or comprising a fragment thereof of at least 10 amino acids, said cDNA being obtainable by the method of Claim 24.
- 26. The cDNA of Claim 25, wherein said cDNA comprises the full protein coding sequence partially included in of one of the sequences of SEQ ID NOs: 38-270.
- 27. A method of making a protein comprising one of the sequences of SEQ ID NO: 271-503, comprising the steps of:

obtaining a cDNA encoding the full protein sequence partially included in one of the sequences of sequence of SEQ ID NO: 38-270;

inserting said cDNA in an expression vector such that said cDNA is operably linked to a promoter;

introducing said expression vector into a host cell whereby said host cell produces the protein encoded by said cDNA; and

isolating said protein.

- 28. An isolated protein obtainable by the method of Claim 27.
- 29. A method of obtaining a promoter DNA comprising the steps of:

obtaining DNAs located upstream of the nucleic acids of SEQ ID NO: 38-270 or the sequences complementary thereto;

screening said upstream DNAs to identify a promoter capable of directing transcription initiation; and

isolating said DNA comprising said identified promoter.

- 30. The method of Claim 29, wherein said obtaining step comprises chromosome walking from said nucleic acids of SEQ ID NO: 38-270 or sequences complementary thereto.
- 31. The method of Claim 30, wherein said screening step comprises inserting said upstream sequences into a promoter reporter vector.
- 32. The method of Claim 30, wherein said screening step comprises identifying motifs in said upstream DNAs which are transcription factor binding sites or transcription start sites.
- 30 An isolated promoter obtainable by the method of Claim 32.

- 34. An isolated or purified protein comprising one of the sequences of SEQ ID NO: 271-503.
- 35. In an array of discrete ESTs or fragments thereof of at least 15 nucleotides in length, the improvement comprising inclusion in said array of at least one of the sequences of SEQ ID NOs: 38-270, or one of the sequences complementary to the sequences of SEQ ID NOs: 38-270, or a fragment thereof of at least 15 consecutive nucleotides.
- 36. The array of Claim 35 including therein at least two of the sequences of SEQ ID NOs: 38-270, the sequences complementary to the sequences of SEQ ID NOs: 38-270, or fragments thereof of at least 15 consecutive nucleotides.
- The array of Claim 35 including therein at least five of the sequences of SEQ ID NOs: 38-270, the sequences complementary to the sequences of SEQ ID NOs: 38-270, or fragments thereof of at least 15 consecutive nucleotides.

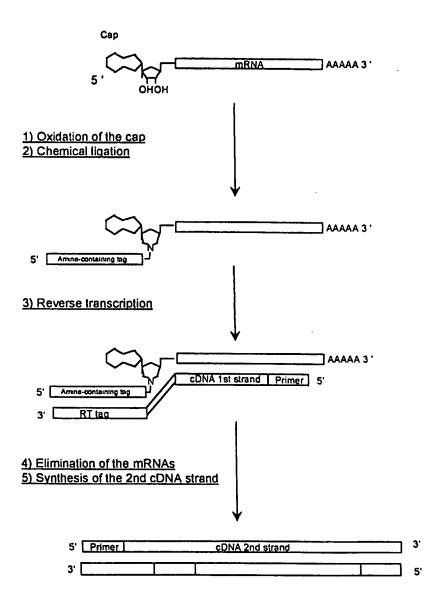


Figure 1

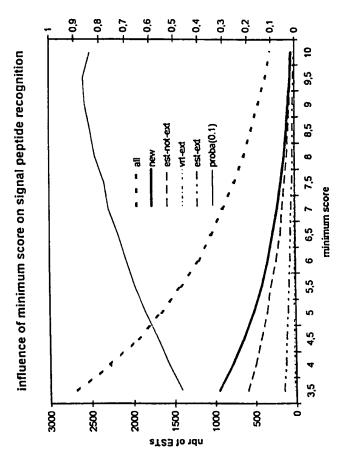


Figure 2

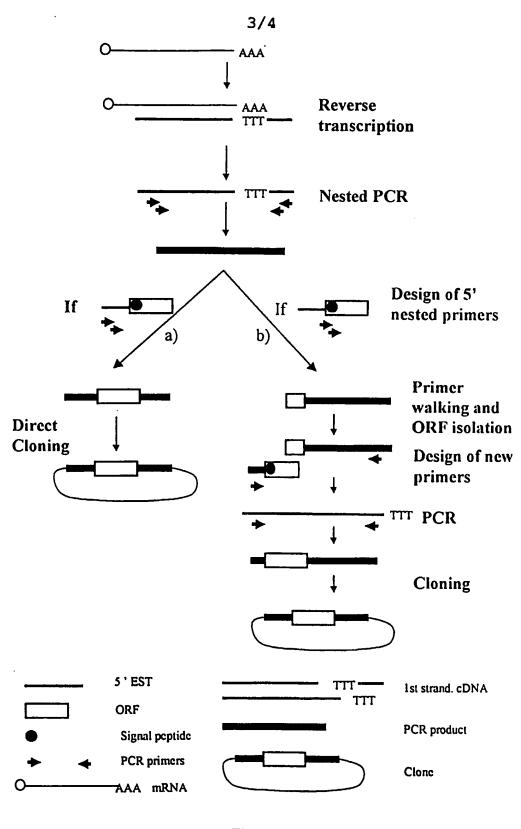
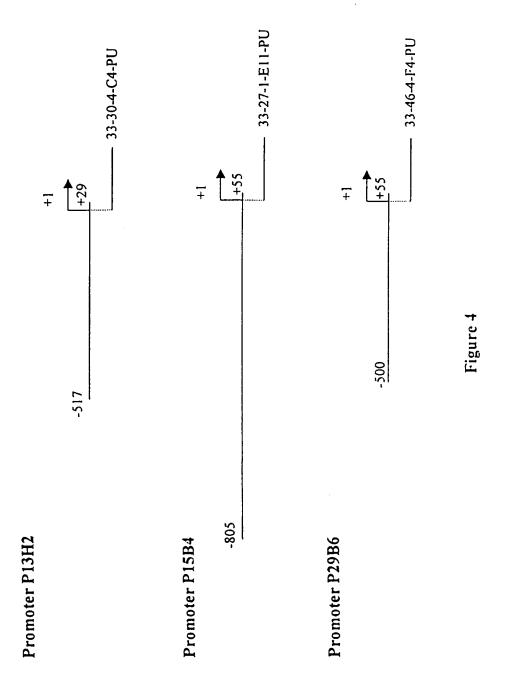


Figure 3



#### SEQUENCE LISTING

- (1) GENERAL INFORMATION:
  - (i) APPLICANT:
    - (A) NAME : GENSET SA
    - (B) STREET :24, RUE ROYALE
    - (C) CITY: PARIS
    - (E) COUNTRY : FRANCE
    - (F) POSTAL CODE (ZIP): 75008
  - (ii) TITLE OF INVENTION: 5' EST FOR SECRETED PROTEINS EXPRESSED IN BRAIN
  - (iii) NUMBER OF SEQUENCES: 503
  - (v) COMPUTER READABLE FORM:
    - (A) MEDIUM TYPE: Floppy Disk
    - (B) COMPUTER: IBM PC compatible
    - (C) OPERATING SYSTEM: Win95
    - (D) SOFTWARE: Word
- (2) INFORMATION FOR SEQ ID NO: 1:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 47 base pairs
    - (B) TYPE: NUCLEIC ACID
    - (C) STRANDEDNESS: SINGLE
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: Other nucleic acid
  - (ix) FEATURE:
    - (A) NAME/KEY: Cap
    - (B) LOCATION: 1
    - (D) OTHER INFORMATION: m7Gppp added to 1
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 1:

GGCAUCCUAC UCCCAUCCAA UUCCACCCUA ACUCCUCCCA UCUCCAC

- (2) INFORMATION FOR SEQ ID NO: 2:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 46 base pairs
    - (B) TYPE: NUCLEIC ACID
    - (C) STRANDEDNESS: SINGLE
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: Other nucleic acid
  - (Mi) SEQUENCE DESCRIPTION: SEQ ID NO: 2:

(2)	INFORMATION FOR SEQ ID NO: 3:	
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 25 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: SINGLE  (D) TOPOLOGY: LINEAR	
	(ii) MOLECULE TYPE: Other nucleic acid	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 3:	
ATCA	AAGAATT CGCACGAGAC CATTA	25
(2)	INFORMATION FOR SEQ ID NO: 4:	
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 25 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: SINGLE  (D) TOPOLOGY: LINEAR	
	(ii) MOLECULE TYPE: Other nucleic acid	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 4:	
TAAT	TGGTCTC GTGCGAATTC TTGAT	25
(2)	INFORMATION FOR SEQ ID NO: 5:	
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 25 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: SINGLE  (D) TOPOLOGY: LINEAR	
	(ii) MOLECULE TYPE: Other nucleic acid	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:	
CCGF	ACAAGAC CAACGTCAAG GCCGC	25
(2)	INFORMATION FOR SEQ ID NO: 6:	
	<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 25 base pairs</li> <li>(B) TYPE: NUCLEIC ACID</li> <li>(C) STRANDEDNESS: SINGLE</li> <li>(D) TOPOLOGY: LINEAR</li> </ul>	

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(ii) MOLECULE TYPE: Other nucleic acid	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6:	
TCACCAGCAG GCAGTGGCTT AGGAG	25
(2) INFORMATION FOR SEQ ID NO: 7:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 25 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: SINGLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: Other nucleic acid	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 7:	
AGTGATTCCT GCTACTTTGG ATGGC	25
(2) INFORMATION FOR SEQ ID NO: 8:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 25 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: SINGLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: Other nucleic acid	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 8:	
GCTTGGTCTT GTTCTGGAGT TTAGA	25
(2) INFORMATION FOR SEQ ID NO: 9:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 25 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: SINGLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: Other nucleic acid	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 9:	

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TCCAGAATGG GAGACAAGCC AATTT

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(2) INFORMATION FOR SEQ ID NO: 10:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 25 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: SINGLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: Other nucleic acid	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 10:	
AGGGAGGAGG AAACAGCGTG AGTCC	25
(2) INFORMATION FOR SEQ ID NO: 11:	
<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 25 base pairs</li> <li>(B) TYPE: NUCLEIC ACID</li> <li>(C) STRANDEDNESS: SINGLE</li> <li>(D) TOPOLOGY: LINEAR</li> </ul>	
(ii) MOLECULE TYPE: Other nucleic acid	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 11:	
ATGGGAAAGG AAAAGACTCA TATCA	25
(2) INFORMATION FOR SEQ ID NO: 12:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 25 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: SINGLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: Other nucleic acid	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 12:	
AGCAGCAACA ATCAGGACAG CACAG	25
(2) INFORMATION FOR SEQ ID NO: 13:	
<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 25 base pairs</li><li>(B) TYPE: NUCLEIC ACID</li><li>(C) STRANDEDNESS: SINGLE</li><li>(D) TOPOLOGY: LINEAR</li></ul>	
(ii) MOLECULE TYPE: Other nucleic acid	

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 13:

ATCAAGAATT CGCACGAGAC CATTA 25

- (2) INFORMATION FOR SEQ ID NO: 14:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 67 base pairs
    - (B) TYPE: NUCLEIC ACID
    - (C) STRANDEDNESS: SINGLE
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: Other nucleic acid
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 14:

ATCGTTGAGA CTCGTACCAG CAGAGTCACG AGAGAGACTA CACGGTACTG GTTTTTTTTT 60

TTTTTVN 67

- (2) INFORMATION FOR SEQ ID NO: 15:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 29 base pairs
    - (B) TYPE: NUCLEIC ACID
    - (C) STRANDEDNESS: SINGLE
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: Other nucleic acid
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 15:

CCAGCAGAGT CACGAGAGAG ACTACACGG 29

- (2) INFORMATION FOR SEQ ID NO: 16:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 25 base pairs
    - (B) TYPE: NUCLEIC ACID
    - (C) STRANDEDNESS: SINGLE
    - (D) TOPOLOGY: LINEAR
    - (ii) MOLECULE TYPE: Other nucleic acid
    - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 16:

CACGAGAGA ACTACACGGT ACTGG 25

(2) INFORMATION FOR SEQ ID NO: 17:

```
(i) SEQUENCE CHARACTERISTICS:
```

- (A) LENGTH: 526 base pairs
- (B) TYPE: NUCLEIC ACID
- (C) STRANDEDNESS: DOUBLE
- (D) TOPOLOGY: LINEAR

### (ii) MOLECULE TYPE: CDNA

### (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo Sapiens
- (F) TISSUE TYPE: Lymph ganglia

### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement (261..376)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 96

region 166..281

id N70479

est

### (ix) FEATURE:

- (A) NAME/KEY: other
- (3) LOCATION: complement (380..486)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 97

region 54..160

id N70479

est

### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement(110..145)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 94

region 403..438

id N70479

est

# (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement(196..229)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 94

region 315..348

id N70479

# (ix) FEATURE:

- (A) NAME/KEY: sig peptide
- (B) LOCATION: 90..140
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 8.2

seq LLLITAILAVAVG/FP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 17:

WO 99/06	552					7	7					PCT/IB9	8/01236
GAGAGAAAGA	. ACTGA	CTGAR	ACGT	rtgag							CTG Leu		113
ACA GCC AT Thr Ala Il	e Leu												161
GAA CGA GA Glu Arg Gl 10	u Lys												209
WTT TTT GT Xaa Phe Va 25													257
CCA TTT CC Pro Phe Pr 40		Phe P											305
CCT GAA TC Pro Glu Se	r Ala									TAAT	ACAAF	<b>RAA</b>	354
gga <b>aa</b> agtca	CRATA	AACCT	GGTC	ACCTG	A AA1	TGA	TTA	GAGO	CCACT	TC C	CTTG	ARAAT	414
СААААТТССТ	GTTAA	TAAAA	RAAA	AACAA	A TG	TAAT	rgaa	ATAC	GCAC	ACA (	GCATI	CTCTA	474
GTCAATATCT	TTAGT	GATCT	TCTT	raata.	A ACI	atga <i>i</i>	AAGC	AAA	VAAAJ	AAA /	AΑ		526

# (2) INFORMATION FOR SEQ ID NO: 18:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 17 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
    (B) LOCATION: 1..17

  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 8.2

seq LLLITAILAVAVG/FP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 18:

Met Lys Lys Val Leu Leu Ile Thr Ala Ile Leu Ala Val Ala Val 10 15

# (2) INFORMATION FOR SEQ ID NO: 19:

### (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 822 base pairs
- (B) TYPE: NUCLEIC ACID
- (C) STRANDEDNESS: DOUBLE
- (D) TOPOLOGY: LINEAR

# (ii) MOLECULE TYPE: CDNA

# (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo Sapiens
- (D) DEVELOPMENTAL STAGE: Fetal
- (F) TISSUE TYPE: kidney

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 260..464
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 96

region 153..357

id H57434

est

### (ix) FEATURE:

- (A) NAME/KEY: other
  (B) LOCATION: 118..184
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 98

region 98..164

id H57434

est

### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 56..113
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 98

region 35..92

id H57434

est

### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 454..485
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 100

region 348..379

id H57434

est

# (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 118..545
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 98

region 1..428

id N27248

est

# (ix) FEATURE:

WO 99/065	552				9			PCT	T/IB98/012
	(B) (C)	LOCATI	EY: other ON: 6536! FICATION MI INFORMATION	ETHOI N:	D: blast identity region 4 id H9477 est	98 1345			
(ix)	(B) (C)	NAME/K LOCATION IDENTI	EY: other ON: 61399 FICATION MI INFORMATION	ETHO		99 344			
(ix)	(B) (C)	NAME/K LOCATION IDENTI	EY: other ON: 4084! FICATION MI INFORMATION	ETHO	D: blast identity region 3 id H0988 est	92 55405			
(ix)	(B) (C)	NAME/K LOCATI IDENTI	EY: other ON: 6039 FICATION M INFORMATION	ETHO		97 6395			
(ix)	(B) (C)	NAME/K LOCATI IDENTI	EY: other ON: 3934 FICATION M INFORMATIO	ETHO N:	D: blast identity region 3 id H2935 est	90 91430			
(ix)	(B) (C)	NAME/K LOCATI IDENTI	EY: sig_pe ON: 3464 FICATION M INFORMATIO	08 ETHO N:	D: Von H score 5.				
(×;)	SEQU	ENCE DE	SCRIPTION:	SEQ	ID NO:	19:			
ACTCCTTTTA	GCAT	AGGGGC	TTCGGCGCCA	GCG	GCCAGCG	CTAGTCGGT	C TGGI	AAGTGC	60
CTGATGCCGA	GTTC	CGTCTC	TCGCGTCTTT	TCC	TGGTCCC	AGGCAAAGC	G GASO	NAGATC	120

CTCAAACGGC CTAGTGCTTC GCGCTTCCGG AGAAAATCAG CGGTCTAATT AATTCCTCTG

GTTTGTTGAA GCAGTTACCA AGAATCTTCA ACCCTTTCCC ACAAAAGCTA ATTGAGTACA

180

240

CGTTCCTGTT GAGTACACGT TCCTGT	TGAT TTACAAAAGG	TGCAGGTATG AGCAGGTCTG	300
AAGACTAACA TTTTGTGAAG TTGTAA	AACA GAAAACCTGT	TAGAA ATG TGG TGG TTT  Met Trp Trp Phe  -20	357
CAG CAA GGC CTC AGT TTC CTT Gln Gln Gly Leu Ser Phe Leu -15			405
GCT GCT TTC ATA TTT TCA TAC Ala Ala Phe Ile Phe Ser Tyr 1, 5			453
GAC CCG GCT TTA CCT TAT ATC Asp Pro Ala Leu Pro Tyr Ile 20			501
AAA TGC TTA TTT GGG GCA ATG Lys Cys Leu Phe Gly Ala Met 35			549
AAA TAGAAATCAG GAARATAATT CA Lys	ACTTAAAG AAKTTC	ATTT CATGACCAAA	602
CTCTTCARAA ACATGTCTTT ACAAGC	ATAT CTCTTGTATT	GCTTTCTACA CTGTTGAATT	662
GTCTGGCAAT ATTTCTGCAG TGGAAA	ATTT GATTTARMTA	GTTCTTGACT GATAAATATG	722
GTAAGGTGGG CTTTTCCCCC TGTGTA	ATTG GCTACTATGT	CTTACTGAGC CAAGTTGTAW	782
TTTGAAATAA AATGATATGA GAGTGA	CACA AAAAAAAAAA		822

# (2) INFORMATION FOR SEQ ID NO: 20:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 21 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION:  $1..\overline{21}$ 

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 5.5

seq SFLPSALVIWTSA/AF

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 20:

Met Trp Trp Phe Gln Gln Gly Leu Ser Phe Leu Pro Ser Ala Leu Val 1 5 10 Ile Trp Thr Ser Ala 20

Pro Asp Asn

(2) INFORMATION FOR SEQ ID NO: 21:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 405 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: CDNA	
<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F) TISSUE TYPE: Testis</pre>	
(ix) FEATURE:  (A) NAME/KEY: other  (B) LOCATION: complement(103398)  (C) IDENTIFICATION METHOD: blastn  (D) OTHER INFORMATION: identity 96  region 1296  id AA442893  est	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 185295     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 5.9</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 21:	
ATCACCTTCT TCTCCATCCT TSTCTGGGCC AGTCCCCARC CCAGTCCCTC TCCTGACCTG 60	)
CCCAGCCCAA GTCAGCCTTC AGCACGCGCT TTTCTGCACA CAGATATTCC AGGCCTACCT 120	)
GGCATTCCAG GACCTCCGMA ATGATGCTCC AGTCCCTTAC AAGCGCTTCC TGGATGAGGG 180	)
TGGC ATG GTG CTG ACC ACC CTC CCC TTG CCC TCT GCC AAC AGC CCT GTG  Met Val Leu Thr Thr Leu Pro Leu Pro Ser Ala Asn Ser Pro Val  -35  -30  -25	Э
AAC ATG CCC ACC ACT GGC CCC AAC AGC CTG AGT TAT GCT AGC TCT GCC  Asn Met Pro Thr Thr Gly Pro Asn Ser Leu Ser Tyr Ala Ser Ser Ala  -20  -15  -10	7
CTG TCC CCC TGT CTG ACC GCT CCA AAK TCC CCC CGG CTT GCT ATG ATG  Leu Ser Pro Cys Leu Thr Ala Pro Xaa Ser Pro Arg Leu Ala Met Met  -5 10	5

CCT GAC AAC TAAATATCCT TATCCAAATC AATAAARWRA RAATCCTCCC TCCARAAGGG 384

405

(2) INFORMATION FOR SEQ ID NO: 22:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 37 amino acids
- (B) TYPE: AMINO ACID
- (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 1..37
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5.9

seq LSYASSALSPCLT/AP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 22:

Met Val Leu Thr Thr Leu Pro Leu Pro Ser Ala Asn Ser Pro Val Asn 1 5 10 15

Met Pro Thr Thr Gly Pro Asn Ser Leu Ser Tyr Ala Ser Ser Ala Leu 20 25 30

Ser Pro Cys Leu Thr 35

- (2) INFORMATION FOR SEQ ID NO: 23:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 496 base pairs
    - (B) TYPE: NUCLEIC ACID
    - (C) STRANDEDNESS: DOUBLE
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: CDNA
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Cancerous prostate
  - (ix) FEATURE:
    - (A) NAME/KEY: other
    - (B) LOCATION: 149..331
    - (C) IDENTIFICATION METHOD: blastn
    - (D) OTHER INFORMATION: identity 98 region 1..183 id AA397994

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 328..485 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 96 region 179..336 id AA397994 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: complement (182..496) (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 97 region 14..328 id AA399680 est (ix) FEATURE: (A) NAME/KEY: sig\_peptide (B) LOCATION: 196..240 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 5.5 seq ILSTVTALTFAXA/LD (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 23: AAAAAATTGG TCCCAGTTTT CACCCTGCCG CAGGGCTGGC TGGGGAGGGC AGCGGTTTAG 60 ATTAGCCGTG GCCTAGGCCG TTTAACGGGG TGACACGAGC NTGCAGGGCC GAGTCCAAGG 120 CCCGGAGATA GGACCAACCG TCAGGAATGC GAGGAATGTT TTTCTTCGGA CTCTATCGAG 180 GCACACAGAC AGACC ATG GGG ATT CTG TCT ACA GTG ACA GCC TTA ACA TTT 231 Met Gly Ile Leu Ser Thr Val Thr Ala Leu Thr Phe -10 GCC ARA GCC CTG GAC GGC TGC AGA AAT GGC ATT GCC CAC CCT GCA AGT 279 Ala Xaa Ala Leu Asp Gly Cys Arg Asn Gly Ile Ala His Pro Ala Ser GAG AAG CAC AGA CTC GAG AAA TGT AGG GAA CTC GAG ASC ASC CAC TCG 327 Glu Lys His Arg Leu Glu Lys Cys Arg Glu Leu Glu Xaa Xaa His Ser GCC CCA GGA TCA ACC CAS CAC CGA AGA AAA ACA ACC AGA AGA AAT TAT 375 Ala Pro Gly Ser Thr Xaa His Arg Arg Lys Thr Thr Arg Arg Asn Tyr TCT TCA GCC TGAAATGAAK CCGGGATCAA ATGGTTGCTG ATCARAGCCC ATATTTAAAT 434 Ser Ser Ala TGGAAAAGTC AAATTGASCA TTATTAAATA AAGCTTGTTT AATATGTCTC AAACAAAAAA 494 P.F. 496

(2) INFORMATION FOR SEQ ID NO: 24:

(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 15 amino acids  (B) TYPE: AMINO ACID  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: PROTEIN	
<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens</pre>	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 115     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 5.5     seq ILSTVTALTFAXA/LD</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 24:	
Met Gly Ile Leu Ser Thr Val Thr Ala Leu Thr Phe Ala Xaa Ala 1 5 10 15	
(2) INFORMATION FOR SEQ ID NO: 25:	
<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 623 base pairs</li><li>(B) TYPE: NUCLEIC ACID</li><li>(C) STRANDEDNESS: DOUBLE</li><li>(D) TOPOLOGY: LINEAR</li></ul>	
(ii) MOLECULE TYPE: CDNA	
<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F) TISSUE TYPE: Testis</pre>	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 4996     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 10.1</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 25:	
AAAGATCCCT GCAGCCCGGC AGGAGAGAAG GCTGAGCCTT CTGGCGTC ATG GAG AGG Met Glu Arg -15	57
CTC GTC CTA ACC CTG TGC ACC CTC CCG CTG GCT GTG GCG TCT GCT GGC Leu Val Leu Thr Leu Cys Thr Leu Pro Leu Ala Val Ala Ser Ala Gly -10	105
TGC GCC ACG ACG CCA GCT CGC AAC CTG AGC TGC TAC CAG TGC TTC AAG Cys Ala Thr Thr Pro Ala Arg Asn Leu Ser Cys Tyr Gln Cys Phe Lys 5 10 15	153

									13	•						
GTC Val 20	AGC Ser	AGC Ser	TGG Trp	ACG Thr	GAG Glu 25	TGC Cys	CCG Pro	CCC Pro	ACC Thr	TGG Trp 30	TGC Cys	AGC Ser	CCG Pro	CTG Leu	GAC Asp 35	201
CAA Gln	GTC Val	TGC Cys	ATC Ile	TCC Ser 40	AAC Asn	GAG Glu	GTG Val	GTC Val	GTC Val 45	TCT Ser	TTT Phe	AAA Lys	TGG Trp	AGT Ser 50	GTA Val	249
						CGC Arg										297
						CCG Pro										345
AGG Arg	CGC Arg 85	TGC Cys	TGT Cys	TCC Ser	TGG Trp	GCT Ala 90	CTC Leu	TGC Cys	AAC Asn	AGG Arg	GCA Ala 95	CTG Leu	ACC Thr	CCA Pro	CAG Gln	393
GAG Glu 100	GGG Gly	CGC Arg	TGG Trp	GCC Ala	CTG Leu 105	CRA Xaa	GGG Gly	GGG Gly	CTC Leu	CTG Leu 110	CTC Leu	CAG Gln	GAC Asp	CCT Pro	TCG Ser 115	441
AGG Arg	GGC Gly	ARA Xaa	AAA Lys	ACC Thr 120	TGG Trp	GTG Val	CGG Arg	CCA Pro	CAG Gln 125	CTG Leu	GGG Gly	CTC Leu	CCA Pro	CTC Leu 130	TGC Cys	489
						CTC Leu										534
TAAC	ACTO	TG G	GTGC	cccc	A CC	TGT	CATI	GGC	SACCA	CRA	CTTC	CACCO	TC 1	TGG	RACAA	594
TAAP	CTCI	CA I	GCCC	CCAA	A AA	AAAA	AAA									623

# (2) INFORMATION FOR SEQ ID NO: 26:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 16 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide(B) LOCATION: 1..16

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 10.1

seq LVLTLCTLPLAVA/SA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 26:

Met Glu Arg Leu Val Leu Thr Leu Cys Thr Leu Pro Leu Ala Val Ala

WO 99/06552 PCT/IB98/01236

1	5	10	15

(2)	INFORMATION	FOR	SEQ	ID	NO:	27:

# (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 848 base pairs
- (B) TYPE: NUCLEIC ACID
- (C) STRANDEDNESS: DOUBLE
- (D) TOPOLOGY: LINEAR

# (ii) MOLECULE TYPE: CDNA

### (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo Sapiens
- (D) DEVELOPMENTAL STAGE: Fetal
- (F) TISSUE TYPE: kidney

# (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 32..73
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 10.7

seq LWLLFFLVTAIHA/EL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 27:

AACT	TTGC	CT 1	rgtgi	TTTC	C AC	CCTG	AAAG			CTG ( Leu 1				55
									-	 GCA Ala	 -			103
										GAT Asp				151
										 ATG Met	 		-	199
				_						 ATT Ile 55	 		-	247
										 TTT Phe	 		-	295
										GTG Val				343
									Ala	TTT Phe		As		391

															ATG Met		39
--	--	--	--	--	--	--	--	--	--	--	--	--	--	--	------------	--	----

GAC CCA TCT GTG CCC ATC TGG ATT ATT ATA TTT GGT GTG ATA TTT TGC
Asp Pro Ser Val Pro Ile Trp Ile Ile Ile Phe Gly Val Ile Phe Cys
125
130
135

ATC ATC ATA GTT GCA ATT GCA CTA CTG ATT TTA TCA GGG ATC TGG CAA

1le lle lle Val Ala lle Ala Leu Leu lle Leu Ser Gly lle Trp Gln

140

145

150

CGT ADA ARA AAG AAC AAA GAA CCA TCT GAA GTG GAT GAC GCT GAA RAT
Arg Xaa Xaa Lys Asn Lys Glu Pro Ser Glu Val Asp Asp Ala Glu Xaa
155 160 170

AAK TGT GAA AAC ATG ATC ACA ATT GAA AAT GGC ATC CCC TCT GAT CCC
Xaa Cys Glu Asn Met Ile Thr Ile Glu Asn Gly Ile Pro Ser Asp Pro
175
180
185

CTG GAC ATG AAG GGA GGG CAT ATT AAT GAT GCC TTC ATG ACA GAG GAT
Leu Asp Met Lys Gly Gly His Ile Asn Asp Ala Phe Met Thr Glu Asp
190 195 200

GAG AGG CTC ACC CCT CTC TGAAGGGCTG TTGTTCTGCT TCCTCAARAA 727
Glu Arg Leu Thr Pro Leu
205

ATTAAACATT TGTTTCTGTG TGACTGCTGA GCATCCTGAA ATACCAAGAG CAGATCATAT 787

WTTTTGTTTC ACCATTCTTC TTTTGTAATA AATTTTGAAT GTGCTTGAAA AAAAAAAAA 847

C 848

# (2) INFORMATION FOR SEQ ID NO: 28:

### (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 14 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(ix) FEATURE:

(A) NAME/KEY: sig peptide

(3) LOCATION: 1..14

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 10.7

seq LWLLFFLVTAIHA/EL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 28:

Met Leu Trp Leu Leu Phe Phe Leu Val Thr Ala Ile His Ala

18 (2) INFORMATION FOR SEQ ID NO: 29: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: SINGLE (D) TOPOLOGY: LINEAR (ii) MOLECULE TYPE: Other nucleic acid (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 29: GGGAAGATGG AGATAGTATT GCCTG 25 (2) INFORMATION FOR SEQ ID NO: 30: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: SINGLE (D) TOPOLOGY: LINEAR (ii) MOLECULE TYPE: Other nucleic acid (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 30: CTGCCATGTA CATGATAGAG AGATTC 26 (2) INFORMATION FOR SEQ ID NO: 31: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 546 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR (ii) MOLECULE TYPE: Genomic DNA (ix) FEATURE: (A) NAME/KEY: promoter (B) LOCATION: 1..517 (ix) FEATURE: (A) NAME/KEY: transcription start site (E) LOCATION: 518 (ix) FEATURE: (A) NAME/KEY: TF binding-site (B) LOCATION: 17..25

(C) IDENTIFICATION METHOD: matinspector prediction

score  $0.9\overline{8}3$ 

sequence TGTCAGTTG

(D) OTHER INFORMATION: name CMYB 01

19 (ix) FEATURE: (A) NAME/KEY: TF binding-site (B) LOCATION: complement (18..27) (C) IDENTIFICATION METHOD: matinspector prediction (D) OTHER INFORMATION: name MYOD Q6 score 0.961 sequence CCCAACTGAC (ix) FEATURE: (A) NAME/KEY: TF binding-site (B) LOCATION: complement (75..85) (C) IDENTIFICATION METHOD: matinspector prediction (D) OTHER INFORMATION: name S8\_01 score 0.960 sequence AATAGAATTAG (ix) FEATURE: (A) NAME/KEY: TF binding-site (B) LOCATION: 94..104 (C) IDENTIFICATION METHOD: matinspector prediction (D) OTHER INFORMATION: name S8 01 score 0.966 sequence AACTAAATTAG (ix) FEATURE: (A) NAME/KEY: TF binding-site (B) LOCATION: complement (129..139) (C) IDENTIFICATION METHOD: matinspector prediction (D) OTHER INFORMATION: name DELTAEF1 01 score 0.960 sequence GCACACCTCAG (ix) FEATURE: (A) NAME/KEY: TF binding-site (B) LOCATION: complement (155..165) (C) IDENTIFICATION METHOD: matinspector prediction (D) OTHER INFORMATION: name GATA C score 0.964 sequence AGATAAATCCA (ix) FEATURE: (A) NAME/KEY: TF binding-site (B) LOCATION: 170..178 (C) IDENTIFICATION METHOD: matinspector prediction (D) OTHER INFORMATION: name CMYB 01 score 0.958 sequence CTTCAGTTG (ix) FEATURE: (A) NAME/KEY: TF binding-site (B) LOCATION: 176..189 (C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name GATA1 02 score 0.959

sequence TTGTAGATAGGACA

# (ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: 180..190

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name GATA\_C

score  $0.9\overline{5}3$ 

sequence AGATAGGACAT

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: 284..299

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name TALIALPHAE47 01

score 0.973

sequence CATAACAGATGGTAAG

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: 284..299

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name TAL1BETAE47 01

score 0.983

sequence CATAACAGATGGTAAG

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: 284..299

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name TAL1BETAITF2\_01

score 0.978

sequence CATAACAGATGGTAAG

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: complement (287..296)

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name MYOD\_Q6

score 0.954
sequence ACCATCTGTT

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(3) LOCATION: complement (302..314)

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name GATA1\_04

score 0.953

sequence TCAAGATAAAGTA

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(3) LOCATION: 393..405

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name IK1 01

score 0.963

sequence AGTTGGGAATTCC

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(3) LOCATION: 393..404

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name IK2\_01

score 0.985

sequence AGTTGGGAATTC

(ix) FEATURE:

	(A) NAME/KEY: TF binding-site (B) LOCATION: 396405 (C) IDENTIFICATION METHOD: matinspector prediction (D) OTHER INFORMATION: name CREL_01 score 0.962 sequence TGGGAATTCC	
(ix)	FEATURE:  (A) NAME/KEY: TF binding-site  (B) LOCATION: 423436  (C) IDENTIFICATION METHOD: matinspector prediction  (D) OTHER INFORMATION: name GATA1_02	
(ix)	FEATURE:  (A) NAME/KEY: TF binding-site  (B) LOCATION: complement(478489)  (C) IDENTIFICATION METHOD: matinspector prediction  (D) OTHER INFORMATION: name SRY 02  score 0.951  sequence TAAAACAAAACA	
(ix)	FEATURE:  (A) NAME/KEY: TF binding-site  (B) LOCATION: 486493  (C) IDENTIFICATION METHOD: matinspector prediction  (D) OTHER INFORMATION: name E2F_02  score 0.957  sequence TTTAGCGC	
(ix)	FEATURE:  (A) NAME/KEY: TF binding-site  (B) LOCATION: complement(514521)  (C) IDENTIFICATION METHOD: matinspector prediction  (D) OTHER INFORMATION: name MZF1_01  score 0.975  sequence TGAGGGGA	
(xi)	SEQUENCE DESCRIPTION: SEQ ID NO: 31:	
rgagtgcagt	GTTACATGTC AGTTGGGTTA AGTTTGTTAA TGTCATTCAA ATCTTCTATG	60
TCTTGATTTG	CCTGCTAATT CTATTATTTC TGGAACTAAA TTAGTTTGAT GGTTCTATTA	120
GTTATTGACT	GAGGTGTGCT AATCTCCCAT TATGTGGATT TATCTATTTC TTCAGTTGTA	180
GATAGGACAT	TGATAGATAC ATAAGTACCA GGACAAAAGC AGGGAGATCT TTTTTCCAAA	240
ATCAGGAGAA	AAAAATGACA TCTGGAAAAC CTATAGGGAA AGGCATAACA GATGGTAAGG	300
	TTGAGTAGGA GAGCCTTCCT GTGGCAACGT GGAGAAGGGA AGAGGTCGTA	360
	GTCAGCTCAG TTAGAAGCAG GGAGTTGGGA ATTCCGTTCA TGTGATTTAG	420
	ATGGCAAATG TGGGACTAAG GGTAGTGATC AGAGGGTTAA AATTGTGTGT	480
	CGCTGCTGGG GCATCGCCTT GGGTCCCCTC AAACAGATTC CCATGAATCT	540
CTTCAT		546

```
(2) INFORMATION FOR SEQ ID NO: 32:
      (i) SEQUENCE CHARACTERISTICS:
            (A) LENGTH: 23 base pairs
            (B) TYPE: NUCLEIC ACID
            (C) STRANDEDNESS: SINGLE
            (D) TOPOLOGY: LINEAR
      (ii) MOLECULE TYPE: Other nucleic acid
      (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 32:
GTACCAGGGA CTGTGACCAT TGC
                                                                     23
(2) INFORMATION FOR SEQ ID NO: 33:
      (i) SEQUENCE CHARACTERISTICS:
            (A) LENGTH: 24 base pairs
            (B) TYPE: NUCLEIC ACID
            (C) STRANDEDNESS: SINGLE
            (D) TOPOLOGY: LINEAR
      (ii) MOLECULE TYPE: Other nucleic acid
      (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 33:
CTGTGACCAT TGCTCCCAAG AGAG
                                                                     24
(2) INFORMATION FOR SEQ ID NO: 34:
      (i) SEQUENCE CHARACTERISTICS:
            (A) LENGTH: 861 base pairs
            (B) TYPE: NUCLEIC ACID
            (C) STRANDEDNESS: DOUBLE
            (D) TOPOLOGY: LINEAR
      (ii) MOLECULE TYPE: Genomic DNA
      (ix) FEATURE:
            (A) NAME/KEY: promoter
            (B) LOCATION: 1..806
      (ix) FEATURE:
            (A) NAME/KEY: transcription start site
            (B) LOCATION: 807
      (ix) FEATURE:
            (A) NAME/KEY: TF binding-site
```

(B) LOCATION: complement(60..70)

(D) OTHER INFORMATION: name NFY Q6

(C) IDENTIFICATION METHOD: matinspector prediction

score 0.956

# sequence GGACCAATCAT

#### (ix) FEATURE:

- (A) NAME/KEY: TF binding-site
- (B) LOCATION: 70..77
- (C) IDENTIFICATION METHOD: matinspector prediction
- (D) OTHER INFORMATION: name MZF1\_01 score 0.962 sequence CCTGGGGA

#### (ix) FEATURE:

- (A) NAME/KEY: TF binding-site
- (B) LOCATION: 124..132
- (C) IDENTIFICATION METHOD: matinspector prediction
- (D) OTHER INFORMATION: name CMYB\_01 score 0.994

sequence TGACCGTTG

### (ix) FEATURE:

- (A) NAME/KEY: TF binding-site
- (B) LOCATION: complement(126..134)
- (C) IDENTIFICATION METHOD: matinspector prediction
- (D) OTHER INFORMATION: name VMYB\_02 score 0.985

sequence TCCAACGGT

### (ix) FEATURE:

- (A) NAME/KEY: TF binding-site
- (B) LOCATION: 135..143
- (C) IDENTIFICATION METHOD: matinspector prediction
- (D) OTHER INFORMATION: name STAT\_01 score 0.968 sequence TTCCTGGAA

### (ix) FEATURE:

- (A) NAME/KEY: TF binding-site
- (B) LOCATION: complement(135..143)
- (C) IDENTIFICATION METHOD: matinspector prediction
- (D) OTHER INFORMATION: name STAT\_01 score 0.951 sequence TTCCAGGAA

# (ix) FEATURE:

- (A) NAME/KEY: TF binding-site
- (B) LOCATION: complement(252..259)
- (C) IDENTIFICATION METHOD: matinspector prediction
- (D) OTHER INFORMATION: name MZF1\_01 score 0.956 sequence TTGGGGGA

# (ix) FEATURE:

- (A) NAME/KEY: TF binding-site
- (B) LOCATION: 357..368
- (C) IDENTIFICATION METHOD: matinspector prediction
- (D) OTHER INFORMATION: name IK2\_01
  score 0.965
  sequence GAATGGGATTTC

### (ix) FEATURE:

- (A) NAME/KEY: TF binding-site
- (B) LOCATION: 384..391

24 (C) IDENTIFICATION METHOD: matinspector prediction (D) OTHER INFORMATION: name MZF1 01 score 0.986 sequence AGAGGGGA (ix) FEATURE: (A) NAME/KEY: TF binding-site (B) LOCATION: complement (410..421) (C) IDENTIFICATION METHOD: matinspector prediction (D) OTHER INFORMATION: name SRY 02 score 0.955 sequence GAAAACAAAACA (ix) FEATURE: (A) NAME/KEY: TF binding-site (B) LOCATION: 592..599 (C) IDENTIFICATION METHOD: matinspector prediction (D) OTHER INFORMATION: name MZF1 01 score 0.960 sequence GAAGGGGA (ix) FEATURE: (A) NAME/KEY: TF binding-site (B) LOCATION: 618..627 (C) IDENTIFICATION METHOD: matinspector prediction (D) OTHER INFORMATION: name MYOD Q6 score 0.981 sequence AGCATCTGCC (ix) FEATURE: (A) NAME/KEY: TF binding-site (B) LOCATION: 632..642 (C) IDENTIFICATION METHOD: matinspector prediction (D) OTHER INFORMATION: name DELTAEF1 01 score 0.958 sequence TCCCACCTTCC (ix) FEATURE: (A) NAME/KEY: TF binding-site (B) LOCATION: complement (813..823) (C) IDENTIFICATION METHOD: matinspector prediction (D) OTHER INFORMATION: name S8 01 score 0.992 sequence GAGGCAATTAT (ix) FEATURE: (A) NAME/KEY: TF binding-site (B) LOCATION: complement (824..831) (C) IDENTIFICATION METHOD: matinspector prediction (D) OTHER INFORMATION: name MZF1 01 score 0.986 sequence AGAGGGGA (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 34: TACTATAGGG CACGCGTGGT CGACGGCCGG GCTGTTCTGG AGCAGAGGGC ATGTCAGTAA 60

TGATTGGTCC CTGGGGAAGG TCTGGCTGGC TCCAGCACAG TGAGGCATTT AGGTATCTCT 120

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CTCAGAGGGC	TAGGCACGAG	GGAAGGTCAG	AGGAGAAGGS	AGGSARGGCC	CAGTGAGARG	240
GGAGCATGCC	TTCCCCCAAC	CCTGGCTTSC	YCTTGGYMAM	AGGGCGKTTY	TGGGMACTTR	300
AAYTCAGGGC	CCAASCAGAA	SCACAGGCCC	AKTCNTGGCT	SMAAGCACAA	TAGCCTGAAT	360
GGGATTTCAG	GTTAGNCAGG	GTGAGAGGG	AGGCTCTCTG	GCTTAGTTTT	GTTTTGTTTT	420
CCAAATCAAG	GTAACTTGCT	CCCTTCTGCT	ACGGGCCTTG	GTCTTGGCTT	GTCCTCACCC	480
AGTCGGAACT	CCCTACCACT	TTCAGGAGAG	TGGTTTTAGG	CCCGTGGGGC	TGTTCTGTTC	540
CAAGCAGTGT	GAGAACATGG	CTGGTAGAGG	CTCTAGCTGT	GTGCGGGGCC	TGAAGGGGAG	600
TGGGTTCTCG	CCCAAAGAGC	ATCTGCCCAT	TTCCCACCTT	CCCTTCTCCC	ACCAGAAGCT	660
TGCCTGAGCT	GTTTGGACAA	AAATCCAAAC	CCCACTTGGC	TACTCTGGCC	TGGCTTCAGC	720
TTGGAACCCA	ATACCTAGGC	TTACAGGCCA	TCCTGAGCCA	GGGGCCTCTG	GAAATTCTCT	780
TCCTGATGGT	CCTTTAGGTT	TGGGCACAAA	ATATAATTGC	CTCTCCCCTC	TCCCATTTTC	840
TCTCTTGGGA	GCAATGGTCA	С				861

# (2) INFORMATION FOR SEQ ID NO: 35:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 20 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: SINGLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: Other nucleic acid
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 35:

CTGGGATGGA AGGCACGGTA

20

# (2) INFORMATION FOR SEQ ID NO: 36:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 20 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: SINGLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: Other nucleic acid
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36:

GAGACCACAC AGCTAGACAA

(2) INFORMATION FOR SEQ ID NO: 37:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 555 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: Genomic DNA

(ix) FEATURE:

(A) NAME/KEY: promoter(B) LOCATION: 1..500

(ix) FEATURE:

(A) NAME/KEY: transcription start site

(B) LOCATION: 501

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: 191..206

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name ARNT\_01 score 0.964

sequence GGACTCACGTGCTGCT

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: 193..204

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name NMYC\_01 score 0.965

sequence ACTCACGTGCTG

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: 193..204

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name USF\_01 score 0.985

sequence ACTCACGTGCTG

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: complement (193..204)

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name USF 01 score 0.985

sequence CAGCACGTGAGT

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: complement(193..204)

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name NMYC 01 score 0.956

sequence CAGCACGTGAGT

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: complement (193..204)

. (C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name MYCMAX\_02

score 0.972

sequence CAGCACGTGAGT

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: 195..202

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name USF C

score 0.997

sequence TCACGTGC

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: complement (195..202)

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name USF\_C

score  $0.\overline{9}91$ 

sequence GCACGTGA

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: complement (210..217)

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name MZF1\_01

score 0.968

sequence CATGGGGA

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: 397..410

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name ELK1\_02

score 0.963

sequence CTCTCCGGAAGCCT

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: 400..409

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name CETS1P54 01

score 0.974

sequence TCCGGAAGCC

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: complement (460..470)

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name AP1\_Q4

score 0.963

sequence AGTGACTGAAC

(ix) FEATURE:

(A) NAME/KEY: TF binding-site

(B) LOCATION: complement (460..470)

(C) IDENTIFICATION METHOD: matinspector prediction

(D) OTHER INFORMATION: name AP1FJ\_Q2

score 0.961

sequence AGTGACTGAAC

	(ix)	FEATURE:  (A) NAME/KEY: TF binding-site  (B) LOCATION: 547555  (C) IDENTIFICATION METHOD: matinspector prediction  (D) OTHER INFORMATION: name PADS_C score 1.000 sequence TGTGGTCTC	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO: 37:	
	CTATAGGGCA	CGCKTGGTCG ACGGCCCGGG CTGGTCTGGT CTGTKGTGGA GTCGGGTTGA	60
	AGGACAGCAT	TTGTKACATC TGGTCTACTG CACCTTCCCT CTGCCGTGCA CTTGGCCTTT	120
	KAWAAGCTCA	GCACCGGTGC CCATCACAGG GCCGGCAGCA CACACATCCC ATTACTCAGA	180
	AGGAACTGAC	GGACTCACGT GCTGCTCCGT CCCCATGAGC TCAGTGGACC TGTCTATGTA	240
	GAGCAGTCAG	ACAGTGCCTG GGATAGAGTG AGAGTTCAGC CAGTAAATCC AAGTGATTGT	300
	CATTCCTGTC	TGCATTAGTA ACTCCCAACC TAGATGTGAA AACTTAGTTC TTTCTCATAG	360
	GTTGCTCTGC	CCATGGTCCC ACTGCAGACC CAGGCACTCT CCGGAAGCCT GGAAATCACC	420
	CGTGTCTTCT	GCCTGCTCCC GCTCACATCC CACACTTGTG TTCAGTCACT GAGTTACAGA	480
	TTTTGCCTCC	TCAATTTCTC TTGTCTTAGT CCCATCCTCT GTTCCCCTGG CCAGTTTGTC	540
	TAGCTGTGTG	GTCTC	555
	(ii) (ii) (vi) (ix)	ATION FOR SEQ ID NO: 38:  SEQUENCE CHARACTERISTICS:  (A) LENGTH: 155 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR  MOLECULE TYPE: CDNA  ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain  FEATURE: (A) NAME/KEY: sig_peptide (B) LOCATION: 4586 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 10.8  seq LLLGLCLGLSLC/VG	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO: 38:	•
,	AGTGTCCCGC	CGGGTCCCCG AGCGTCCCGC GCCCTCGCCC CGCC ATG CTC CTG CTG Met Leu Leu Leu	56
		C MCC CDC CCC CMC MCC CMC MCM CMC CCC CC	

Leu -10	Gly	Leu	Cys	Leu	Gly -5	Leu	Ser	Leu	Cys	Val 1	Gly	Ser	Gln	Glu 5	Glu	
						TCT Ser										152
AGG																155

# (2) INFORMATION FOR SEQ ID NO: 39:

Arg

12	1 00000000	CHARACTERISTICS:

- (A) LENGTH: 427 base pairs
- (B) TYPE: NUCLEIC ACID
- (C) STRANDEDNESS: DOUBLE
- (D) TOPOLOGY: LINEAR

# (ii) MOLECULE TYPE: CDNA

### (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo Sapiens
- (F) TISSUE TYPE: Brain

# (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 191..268
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 10.8

seq VLLFFVLLGMSQA/GS

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 39:

AAGTACACGC GGMGAACTGG GAAGACAGAA AGWWCAATCC TTTAAGGGAG AACCTAGAAG	60
CCATTCAACA AGGTTAAAAT CTTTAGGCTT CCGAGGATTT GGTAGACAGA TCAGAGGCAC	120
GTTTCCCACA ACTGCGAAGA GGCGCTGAGG CAATTCTGCA AGAAGATTTT GGGGTTTTGG	180
AAAAGAAGCT ATG GAA AAC GGA GGG GCA GGC ACT CTG CAG ATA AGG CAA Met Glu Asn Gly Gly Ala Gly Thr Leu Gln Ile Arg Gln -25 -20 -15	229
GTC CTG CTT TTC TTT GTT TTG CTG GGA ATG TCT CAG GCG GGC TCT GAA Val Leu Leu Phe Phe Val Leu Leu Gly Met Ser Gln Ala Gly Ser Glu -10 -5 1	277
ACT GGG AAC TTT TTG GTG ATG GAG GAA TTG CAG AGC GGG AGC TTT GTA Thr Gly Asn Phe Leu Val Met Glu Glu Leu Gln Ser Gly Ser Phe Val 5 10 15	325
GGA AAT TTG GCA AAG ACC CTG GGA CTC GAG GTG AGT GAG CTG TCT TCG Gly Asn Leu Ala Lys Thr Leu Gly Leu Glu Val Ser Glu Leu Ser Ser 25 30 35	373
CGG GGG GCT CGG GTG GTT TCT AAT GAT AAC AAA GAG TGT TTG CAG CTG Arg Gly Ala Arg Val Val Ser Asn Asp Asn Lys Glu Cys Leu Gln Leu 40 45 50	421

GAC	ACG		427
Asp	Thr	•	

<i>1</i> ? ` `	INFORMATION	TOD CTO	7.0	N10 -	<b>Λ</b> Λ .
	INFURMATION	PUR SEU	117	INI ) :	44 () :

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 398 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:

-30

-15

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 12..389
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 10

seq LKLLLFLSTELQA/SQ

338

386

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 40:

	G CCG GGT CCC CAA CO 1 Pro Gly Pro Gln P -120	
	GCG CTG GGG TAT AA Ala Leu Gly Tyr Ly -105	
	GCG GCA GAG GGT GG Ala Ala Glu Gly Gl -8	y Leu Ser Ser
	 TGG TTA GGC TCT CA Trp Leu Gly Ser Gl -70	
	ACG TCT GCT GGA AG Thr Ser Ala Gly Ar -55	
Glu Ser Phe Gln	GGC TTT TTA AAA GA Gly Phe Leu Lys Gl -40	

CCA TAT TCT GTA CTC ATA TCA GGA GAT ATT AAA GAT CGT TTA AAA AAG

Pro Tyr Ser Val Leu Ile Ser Gly Asp Ile Lys Asp Arg Leu Lys Lys

AAG GAG GAC TGT TTG AAA CTT CTA TTA TTT TTA AGT ACA GAA CTT CAA

Lys Glu Asp Cys Leu Lys Leu Leu Phe Leu Ser Thr Glu Leu Gln

-10

-25

-5

GCT TCA CAG ATA Ala Ser Gln Ile 1	398
(2) INFORMATION FOR SEQ ID NO: 41:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 201 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: CDNA	
<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F) TISSUE TYPE: Brain</pre>	
(ix) FEATURE:  (A) NAME/KEY: sig_peptide  (B) LOCATION: 70147  (C) IDENTIFICATION METHOD: Von Heijne matrix  (D) OTHER INFORMATION: score 9.6  seq WLIALASWSWALC/RI	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 41:	
AGCCCGGTTT CGTGCCCGCG GCCGACTGCG CASCTGTCCG CGAGTCTGAG ATACTTACAG	60
AGAGCTACA ATG GAA AAG TCC TGG ATG CTG TGG AAC TTT GTT GAA AGA TGG Met Glu Lys Ser Trp Met Leu Trp Asn Phe Val Glu Arg Trp -25 -20 -15	111
CTA ATA GCC TTG GCT TCA TGG TCT TGG GCT CTC TGC CGT ATT TCT CTT Leu Ile Ala Leu Ala Ser Trp Ser Trp Ala Leu Cys Arg Ile Ser Leu -10 -5 1	159
TTA CCT TTA ATA GTG ACT TTT CAT CTG TAT GGA GGT TCG GGG Leu Pro Leu Ile Val Thr Phe His Leu Tyr Gly Gly Ser Gly 5 10 15	201
(2) INFORMATION FOR SEQ ID NO: 42:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 272 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: CDNA	
(vi) CRIGINAL SOURCE:	

(A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain

	( )	lx). E	TEAT													
			(A)	NAME	:/KEY	: si	.g_pe	ptic	de							
(B) LOCATION: 6113																
(C) IDENTIFICATION METHOD: Von Heijne matrix																
(D) OTHER INFORMATION: score 9.5																
									seq	LGLI	LLAF	RHWCI	A/GV	7		
	_						•									
	()	(i) S	EQUE	ENCE	DESC	CRIPT	CION:	SEC	O ID	NO:	42:					
מכמים	ח א יחים	יר כי			-m -c	· · · · ·	- 2 - 21		-m -c							
AGA															CT CAC	
	(*16		111 G. 35	L11 L1	IT M	. g . i i	11 G.		La Va	T A	la G.		La PI 25	ne Se	er His	
		•	, ,					, 0				4	2.5			
TGC	CTG	GGC	TTC	TGT	GGA	ATG	AGA	СТС	GGG	СТС	СТТ	СТА	СТТ	GCA	AGA	98
					Gly											,,
•	-20	•		•		-15					-10					
CAC	TGG	TGC	ATT	GCA	GGT	GTG	TTT	CCG	CAG	AAG	TTT	GAT	GGT	GAC	AGT	146
His	Trp	Cys	Ile	Ala	Gly	Val	Phe	Pro	Gln	Lys	Phe	Asp	Gly	Asp	Ser	
-5					1				5					10		
GCC	TAC	GTG	GGG	ATG	AGT	GAC	GGA	AAC	CCA	GAG	CTC	CTG	TCA	ACC	AGC	194
Ala	Tyr	Val		Met	Ser	Asp	Gly		Pro	Glu	Leu	Leu		Thr	Ser	
			15					20					25			
CAG	ACC	TAC	ממכ	ccc	CAG	N.C.C	GNG	אאר	אאר	CNN	CNC	T A T	CAC	B ITC	ccc	242
					Gln											242
O	••••	30	ASII	OLY	GIII	Jer	35	N311	N311	GIU	vah	40	GIU	116	FIO	
		-										40				
CCG	ATA	AÇA	CCT	CCC	AAC	CTC	CČG	GAA	GCG							272
					Asn											
	45					50										
(2)	INFO	ORMA!	TION	FOR	SEQ	ID I	10: 4	13:								
(i) SPOUENCE CHARACTERISTICS:																

- - (A) LENGTH: 186 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 28..99
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 9.1

seq LVVFLLLPLASGP/QV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 43:

Met Glu Lys Gly Asn Ala Phe Leu Lys

Met Glu Lys Gly Asn Ala Phe Leu Lys -20									
AAT AGG TTA GTT GTT TTT CTC CTG CCC CTA GCA TCA GGA CCA CAG Asn Arg Leu Val Val Phe Leu Leu Pro Leu Ala Ser Gly Pro Gln -15 -10 -5 1	102								
GTA AAA AGG AAA AGC GAA ATT ACG AAA CTT ATA AAG GCC ACG CGA ATC Val Lys Arg Lys Ser Glu Ile Thr Lys Leu Ile Lys Ala Thr Arg Ile 5 10 15	150								
ATT TGT TTA TTC AAT AAA TTT AGT AGA GGA AAC GGG Ile Cys Leu Phe Asn Lys Phe Ser Arg Gly Asn Gly 20 25	186								
(2) INFORMATION FOR SEQ ID NO: 44:  (i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 400 base pairs									
(B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR									
(ii) MOLECULE TYPE: CDNA									
<ul><li>(vi) ORIGINAL SOURCE:</li><li>(A) ORGANISM: Homo Sapiens</li><li>(F) TISSUE TYPE: Brain</li></ul>									
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 164235     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 9</pre>									
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 44:									
AGTTAATTTG AACAAAATAT AGTAAGTATA CTATTATTTT CCATGTCTTT CTAGGCTTTT	60								
TAAACTCTGC AGTGTATTTA CTGTACTCTC GTAGAAGGAG TGCCATCAAC TGCAATTGGT	120								
ACAAATTGTG CTTATTTTTC TGCTCTTTTC ACGTTCCCAA AAT ATG TTC CCA TTC Met Phe Pro Phe	175								
AAC CAG GCA GGT CTT CCT ACT CTC ATG CTC ATT GTT TTT CAT GCT Asn Gln Ala Gly Leu Pro Thr Leu Leu Met Leu Ile Val Phe His Ala -20 -15 -10 -5	223								
GCT TCC ATG GCT TTA CAG AGA CTC TTC CTC TTC GCT TTG GTC TGG CAT Ala Ser Met Ala Leu Gln Arg Leu Phe Leu Phe Ala Leu Val Trp His 1 5 10	271								
TCA AAA CCC TCA GGA CTG ATG ACA GGC AAA CTA GAA TCT CAA ATT CCC Ser Lys Pro Ser Gly Leu Met Thr Gly Lys Leu Glu Ser Gln Ile Pro 15 20 25	319								

CAT GAA AAG TEG ACT CAT ATC TCT GTC ATG CAT GGT CCC CTC AGT TCC 367

His Glu Lys Leu Thr His Ile Ser Val Met His Gly Pro Leu Ser Ser 35 CAT CAC TCA TAC ACT CAC ATA CAT TTA TTT TTA 400 His His Ser Tyr Thr His Ile His Leu Phe Leu (2) INFORMATION FOR SEQ ID NO: 45: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 297 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR (ii) MOLECULE TYPE: CDNA (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain (ix) FEATURE: (A) NAME/KEY: sig peptide (B) LOCATION:  $1..\overline{228}$ (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 8.8 seg SLLLWMSSLPSLG/EK (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 45: ATG ACT TCA CGT AGC TTG CGT CGC TGC TCC TGT CTC CGT GTA ACT CAC 48 Met Thr Ser Arg Ser Leu Arg Arg Cys Ser Cys Leu Arg Val Thr His -70 AAT AAA GAG ATT TTG GCA TCA ACC GTG AGC TTA GGG GTA GAA GGG TAT 96 Asn Lys Glu Ile Leu Ala Ser Thr Val Ser Leu Gly Val Glu Gly Tyr -55 ATG TTA GGA GGT GGG AGC AGA ATC AAT TCT TCA AAT CTT AAT GAT GGT 144 Met Leu Gly Gly Ser Arg Ile Asn Ser Ser Asn Leu Asn Asp Gly

GAA GAA GAG TGC TCA CCA GAT TCC CTT CTG GTC TGG AAA AAG AAA TCC 192 Glu Glu Glu Cys Ser Pro Asp Ser Leu Leu Val Trp Lys Lys Ser -20 CTT CTT TTG TGG ATG TCA TCT CTA CCA TCT CTC GGT GAA AAA TAT TTC 240 Leu Leu Trp Met Ser Ser Leu Pro Ser Leu Gly Glu Lys Tyr Phe AAG AGA ATO CTA AGA TGG AGA GAG CAT TGG AAG TCA TCC GGC CCA ATT 288 Lys Arg Ile Leu Arg Trp Arg Glu His Trp Lys Ser Ser Gly Pro Ile 10 CCC TTG TGG 297 Pro Leu Trp

(2) INFORMATION FOR SEQ ID NO: 46:										
<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 213 base pairs</li> <li>(B) TYPE: NUCLEIC ACID</li> <li>(C) STRANDEDNESS: DOUBLE</li> <li>(D) TOPOLOGY: LINEAR</li> </ul>										
(ii) MOLECULE TYPE: CDNA										
<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F) TISSUE TYPE: Brain</pre>										
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 10168     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 8.8</pre>										
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 46:										
AACTGTAGC ATG TGG ACA GCC AGT GCC ATG GAT TTC AGA ACC TGC ATT GCC 51  Met Trp Thr Ala Ser Ala Met Asp Phe Arg Thr Cys Ile Ala  -50 -45 -40										
AGT KGA CTG CCT GCT TTG TGC TAC GTG CAG GCC TGC CGC GCC CTG ATG Ser Xaa Leu Pro Ala Leu Cys Tyr Val Gln Ala Cys Arg Ala Leu Met -35	99									
ATT GCT GCC TCG GTC CTG GGT CTG CCG GCC ATT TTA CTG CTG CTG ACT Ile Ala Ala Ser Val Leu Gly Leu Pro Ala Ile Leu Leu Leu Thr -20 -15 -10	147									
GTT CTT CCC TGC ATC SGG ATG GGC CAG GAG CCC GGT GTG GCT AAG TAC Val Leu Pro Cys Ile Xaa Met Gly Gln Glu Pro Gly Val Ala Lys Tyr -5 1 5	195									
AGG SGG GCC CAG CTG GCT Arg Xaa Ala Gln Leu Ala 10 15	213									
(2) INFORMATION FOR SEQ ID NO: 47:										
<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 319 base pairs</li> <li>(B) TYPE: NUCLEIC ACID</li> <li>(C) STRANDEDNESS: DOUBLE</li> <li>(D) TOPOLOGY: LINEAR</li> </ul>										
(11) MOLECULE TYPE: CDNA										

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain

O 99/06552	36	PCT/IB98/01236
(ix).FEAT	JRE:	
(A)	NAME/KEY: sig peptide	
(B)	LOCATION: 62196	
(C)	IDENTIFICATION METHOD: Von He	ijne matrix

seq FALLSLSHPTCQA/GA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 47:

(D) OTHER INFORMATION: score 8.5

ATTGGTGCAG AGGCCCTTCT TGTCTCCACA CCAGAAGGAG CTGAGCAGAG GGGCCACAGC	60
G ATG GGA CCC CCT CCA ACC CAC ATT AAA TAC CTC CAC CTG AAT ATT TAT Met Gly Pro Pro Pro Thr His Ile Lys Tyr Leu His Leu Asn Ile Tyr -45 -35 -30	109
TGC AAC GGC AAG AGC ACT GCA CCT GGA ATC CGG TCT CAC AGC CTT GGA Cys Asn Gly Lys Ser Thr Ala Pro Gly Ile Arg Ser His Ser Leu Gly -25 -20 -15	157
TTT GCC TTG CTA AGC CTC AGT CAT CCA ACC TGC CAG GCA GGT GCA CCT Phe Ala Leu Leu Ser Leu Ser His Pro Thr Cys Gln Ala Gly Ala Pro -10 -5 1	205
GCC GCA GCC CTG CCT TCT CTG TGG AGC TGG TGC TCT CGG GGT GCA CGA Ala Ala Ala Leu Pro Ser Leu Trp Ser Trp Cys Ser Arg Gly Ala Arg 5 10 15	253
GTC AGG GTT GGG AGG ATG CTT TCT CAC CTG TAC ACC TGT GGA TGG TAC Val Arg Val Gly Arg Met Leu Ser His Leu Tyr Thr Cys Gly Trp Tyr 20 25 30 35	301
GAT CAC AAC CCC CAT GGG Asp His Asn Pro His Gly 40	319

# (2) INFORMATION FOR SEQ ID NO: 48:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 260 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 204..251
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 8.5

seq LLTFLAFTTLLFA/PP

(Mi) SEQUENCE DESCRIPTION: SEQ ID NO: 48:

ACTGTTATTC TTCACATATT ATCTCTATTA GTADVWGGCT TCTAGTCACC CAAGTTAGAA	60
ATCTGTGAGT CATCTTTGT TTCTTCCCTT TCCCTTACTG TTTAGTTTTA ATTGCTAAGT	120
CTTGTTAATA CTACATCAGG TATGATTTTA AAAACATTTT TGATGTTCTA CTGCCACCAC	180
CTTAGTTCTG GTACTCATTT TGC ATG TTT TGT CTT TTG ACT TTC CTT GCT TTT  Met Phe Cys Leu Leu Thr Phe Leu Ala Phe  -15 -10	233
ACA ACT CTT CTT TTC GCA CCC CCA TGG Thr Thr Leu Leu Phe Ala Pro Pro Trp -5 1	260
(2) INFORMATION FOR SEQ ID NO: 49:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 365 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA	
<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F) TISSUE TYPE: Brain</pre>	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 126212     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 8.4</pre>	
ACAAATGTGT TATGATTTTC CAGGCCCTTC TTCATCTGCT CTCCCTTCCT TTTGAGCATT	60
ATCCCATTTC ATGCCCCAC ACAGATTCTA GCCATACCCC ATGACTTACA ATTTCCCCAC	120
AAAGA ATG CAC TGT GGC TCC ACT CCA GGA CTT TGC CCA TGC TGG GTC CCC  Met His Cys Gly Ser Thr Pro Gly Leu Cys Pro Cys Trp Val Pro  -25 -20 -15	170
TTC CTG AAA TGC CTT CTA GCT GTT CTC TCT TCC CTG TTT GCT GCC ATT Phe Leu Lys Cys Leu Leu Ala Val Leu Ser Ser Leu Phe Ala Ala Ile -10 -5 1	218
TCC GTG GAC AGA CTA TAC TTG TCT TTC TGT TCT AAT TGC TCT GAA ATA Ser Val Asp Arg Leu Tyr Leu Ser Phe Cys Ser Asn Cys Ser Glu Ile 5 10 15	266
TAC CTC TGG CCC CCC AGC TTT CCT GCT CCC CCA TCC CCT GTA GTC CTT Tyr Leu Trp Pro Pro Ser Phe Pro Ala Pro Pro Ser Pro Val Val Leu 20 25 30	314

CTA GTT TTC CTG TGT CCC CAT GGG ACT TCT TTA TCC TTT TTG AAG CTA Leu Val Phe Leu Cys Pro His Gly Thr Ser Leu Ser Phe Leu Lys Leu 40 CCG 365 Pro (2) INFORMATION FOR SEQ ID NO: 50: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 69 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR (ii) MOLECULE TYPE: CDNA (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain (ix) FEATURE: (A) NAME/KEY: sig\_peptide (B) LOCATION: 1..48 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 8.3 seq VCSALLLLGIVSS/KP (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 50: ATG AAT TTA GTT TGT TCA GCT CTT TTA CTT CTT GGA ATA GTA TCT TCC 48 Met Asn Leu Val Cys Ser Ala Leu Leu Leu Gly Ile Val Ser Ser AAA CCC TAT ATG AGA AAG CGG 69 Lys Pro Tyr Met Arg Lys Arg (2) INFORMATION FOR SEQ ID NO: 51: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 184 base pairs (3) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR (ii) MOLECULE TYPE: CDNA (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain (ix) FEATURE: (A) NAME/KEY: sig\_peptide (3) LOCATION: 44..148 (C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION:	score 8.3 seq AAMLIGLLAWLQT/VP
(xi) SEQUENCE DESCRIPTION: SE	CQ ID NO: 51:
AAATTACAAG AAAGCTGGAC TTGCCGCTGT GC	GTCTCAGGA GAA ATG AGT GTT CTT 55 Met Ser Val Leu -35
GAT GAC AGG CAA AGG GAC ATC TTA GTT Asp Asp Arg Gln Arg Asp Ile Leu Val -30 -25	
CTG GAA GCC GCC ATG TTA ATA GGA TTA Leu Glu Ala Ala Met Leu Ile Gly Leu -15 -10	A CTA GCC TGG CTC CAG ACA GTG 151 Leu Ala Trp Leu Gln Thr Val -5 1
CCT GCT CAT GGC TGC CAG TTC TTA CCC Pro Ala His Gly Cys Gln Phe Leu Pro 5	o Ile Arg
(2) INFORMATION FOR SEQ ID NO: 52:  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 251 base pa (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBL (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapi (F) TISSUE TYPE: Brain  (ix) FEATURE: (A) NAME/KEY: sig_pepti (B) LOCATION: 138197 (C) IDENTIFICATION METH (D) OTHER INFORMATION:	de  OD: Von Heijne matrix score 8.3 seq LLIICHYLPLSLC/IP
AATTTTTTT CACTTTCCTA AAAGCCTTCC CT	TTTGCCCAT GATGCCAATG ACTAGCTCTG 60
TCCTGAAGCA ATAGCTAGTA CTTTCCCTCC TT	CCTGCCAC CTAGCATCCA GCCGAACCTT 120
GRATCAATAC CAGTAAA ATG GGT GTG AAC Met Gly Val Asn -20	GGA AGG AGG CTG CTC ATT ATT  Gly Arg Arg Leu Leu Ile Ile  -15  -10
TGC CAT TAT TTA CCT CTG AGT CTG TGC Cys His Tyr Leu Pro Leu Ser Leu Cys -5	
ANT TOT CTC CCG CGC AAC ACC CCC CC	T GTC AGG 251

Asn Ser Leu Pro Arg Asn Thr Pro Pro Val Arg

(2)	INFOR	MATION	FOR	SEQ	ID	NO:	53:
	(i)	SEQUEN	ICE C	CHARA	CTE	RIS	rics

(A) LENGTH: 154 base pairs (B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 26..118

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 8.2

seq LECLLLYLAESSG/LR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 53:

ACAGAACTAA CCTAGAAAGA ATGAT ATG AAA CTT CGT GAG TGC CCG GCC CTC 52 Met Lys Leu Arg Glu Cys Pro Ala Leu CGA TGG TCC CAG CTG TCC CAG CAC AAG CTG GAG TGT CTA TTG CTT TAC 100 Arg Trp Ser Gln Leu Ser Gln His Lys Leu Glu Cys Leu Leu Leu Tyr -20 -15 CTG GCA GAG AGC TCC GGG CTC AGA ACA GGA AAT GTG GGA GTT CTC CAC 148 Leu Ala Glu Ser Ser Gly Leu Arg Thr Gly Asn Val Gly Val Leu His 1 CCA AGG 154

#### (2) INFORMATION FOR SEQ ID NO: 54:

Pro Arg

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 485 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide(B) LOCATION: 78..404

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 8.2

seq LLRLPQLPPXCSA/GE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 54:

ACC	CTTTC	CGT (	CCGC	гстс	AT TO	GGCT	CTGC'	r GC	AGCC	CTGA	CCA	ACGC!	rcc i	ATA	GCCGG	60
GAT	CCAGO	CCA 1	ract:		ATG (			Arg (					Ala 1			110
					CAT His											158
					GAG Glu											206
					GTG Val											254
					ATT Ile -45											302
					ACT Thr					Asp						350
					CTT Leu											398
					AGC Ser											446
					TGG Trp 20						Pro					485

# (2) INFORMATION FOR SEQ ID NO: 55:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 276 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain	
<ul> <li>(ix) FEATURE:</li> <li>(A) NAME/KEY: sig_peptide</li> <li>(B) LOCATION: 199261</li> <li>(C) IDENTIFICATION METHOD: Von Heijne matrix</li> <li>(D) OTHER INFORMATION: score 8</li> <li>seq LFLVAVLVKVAEA/RK</li> </ul>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 55:	
AGCCTCTCCT GCACCCTCAG CCGGCGCGCT TCTCTTATGG GCGTCTGCTG CAGTCTGGCT	60
GCGGTCGAAC TGAAAGCGGC GGCGGGAGAC CAAACTTAGA CCCCGCTGTG GACTAGAGAA	120
CTCAGAGAAG GCAGAGGGAG AGGGAGAGAG AGASABWBAA GGGACCCGAG GAGGAGGCTT	180
CCATCACGTC ATTGCAGG ATG TTC TGG AAG CTT TCC CTG TCC TTG TTC CTG  Met Phe Trp Lys Leu Ser Leu Phe Leu  -20  -15	231
GTG GCG GTG CTG GTG AAG GTG GCG GAA GCC CGG AAG AAC CGG TCG Val Ala Val Leu Val Lys Val Ala Glu Ala Arg Lys Asn Arg Ser -10 -5 1 5	276
(2) INFORMATION FOR SEQ ID NO: 56:  (i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 197 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: CDNA	
<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F) TISSUE TYPE: Brain</pre>	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 120173     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 7.9</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 56:	
AAGTTTCCCG GGAGAGACGA AAGCAGGAAC GAGAGCGGAG GNAGCACAGT CCGCCGAGCA	60
CAAGCTCCAG CATCCCGTCA GGGGTTGCAG GTGTGTGGGA GGCTTGAAAC TGTTACAAT	119
ATG GCT TTC CTT GGA CTC TTC TCT TTG CTG GTT CTG CAA AGT ATG GCT Met Ala Phe Leu Gly Leu Phe Ser Leu Leu Val Leu Gln Ser Met Ala -15 -10 -5	167
ACA GGG GCC ACT TTC CCT GAG GAA GCC CCG	197

Thr Gly Ala Thr Phe Pro Glu Glu Ala Pro 1 5

INFC	RMAT	CION	FOR	SEQ	ID N	10: 5	57:								
(i	.) SE	(A) (B) (C)	LENG TYPE STRA	TH: : NU .NDEC	299 CLEI NESS	base C AC : DC	pai ID UBLE								
(i	i) M	OLEC	ULE	TYPE	: CD	NA									
(∨	·i) C	(A)	ORGA	NISM	l: Ho		-	ens							
(i	×) F	(A) (B) (C)	NAME LOCA IDEN	TION	: 90 CATI	14 ON M	3 ETHO	D: V	e 7.	9					
(°x	:i) S	EQUE	NCE	DESC	RIPT	'ION:	SEC	) ID	NO:	57:					
AGCG	GA S	STAC	CACA	G TO	CGCC	GAGO	C AC	AAGCI	CCA	GCAT	cccc	STC A	AGGGT	TTGCAG	60
GTGG	GA G	GCTI	GAAF	AC TO	STTAC	 CAAT									113
															161
															209
															257
															299
		13UQB (A) (B)	CE C LENC	CHARA STH: E: NU	ACTES 370 JCLES	RIST: base	ICS: e pa: CID								
	(i) (v) (i) (x) AGCG CTG Leu ATT Ile AGC AATT Ile AGC AII INFO	(ii) SE  (ii) M  (vi) C  (ix) E  (xi) S  AGCGGA S  GTGGGA C  CTG GTT  Leu Val  ATT GCT  Ile Ala  AGC TGG  Ser Trp  25  GCC TTT  Ala Phe  40	(i) SEQUEN (A) (B) (C) (D) (ii) MOLEC (vi) ORIGI (A) (F) (ix) FEATO (A) (B) (C) (D) (xi) SEQUE AGCGGA SSTAC GTGGGA GGCTT CTG GTT CTG Leu Val Leu ATT GCT GAC Ile Ala Asp 10 AGC TGG AGA Ser Trp Arg 25 GCC TTT CTC Ala Phe Leu 40 INFORMATION (i) SEQUEN (A) (B)	(i) SEQUENCE O  (A) LENG (B) TYPE (C) STRA (D) TOPO  (ii) MOLECULE  (vi) ORIGINAL (A) ORGA (F) TISS  (ix) FEATURE: (A) NAME (B) LOCA (C) IDEN (D) OTHE  (xi) SEQUENCE  AGCGGA SSTAGCACA  GTGGGA GGCTTGAAA  CTG GTT CTG CAA Leu Val Leu Gln  ATT GCT GAC TTG Ile Ala Asp Leu 10  AGC TGG AGA AGG Ser Trp Arg Arg 25  GCC TTT CTC CTC Ala Phe Leu Leu 40  INFORMATION FOR (i) SEQUENCE ( (A) LENC (B) TYPE	(i) SEQUENCE CHARA  (A) LENGTH:  (B) TYPE: NU  (C) STRANDED  (D) TOPOLOGY  (ii) MOLECULE TYPE  (vi) ORIGINAL SOUR  (A) ORGANISM  (F) TISSUE T  (ix) FEATURE:  (A) NAME/KEY  (B) LOCATION  (C) IDENTIFI  (D) OTHER IN  (xi) SEQUENCE DESC  AGCGGA SSTAGCACAG TO  CTG GTT CTG CAA AGT  Leu Val Leu Gln Ser  -5  ATT GCT GAC TTG TCA  Ile Ala Asp Leu Ser  10  AGC TGG AGA AGG GAA  Ser Trp Arg Arg Glu  25  GCC TTT CTC CTC CAT  Ala Phe Leu Leu His  40  INFORMATION FOR SEQ  (i) SEQUENCE CHARA  (A) LENGTH:  (B) TYPE: NU	(i) SEQUENCE CHARACTER (A) LENGTH: 299 (B) TYPE: NUCLEI (C) STRANDEDNESS (D) TOPOLOGY: LI  (ii) MOLECULE TYPE: CD  (vi) ORIGINAL SOURCE: (A) ORGANISM: HO (F) TISSUE TYPE:  (ix) FEATURE: (A) NAME/KEY: Si (B) LOCATION: 90 (C) IDENTIFICATI (D) OTHER INFORM  (xi) SEQUENCE DESCRIPT  AGCGGA SSTAGCACAG TCCGCO  STGGGA GGCTTGAAAC TGTTAC  CTG GTT CTG CAA AGT ATG Leu Val Leu Gln Ser Met -5  ATT GCT GAC TTG TCA GTG Ile Ala Asp Leu Ser Val 10  AGC TGG AGA AGG GAA GGA Ser Trp Arg Arg Glu Gly 25  GCC TTT CTC CTC CAT TTA Ala Phe Leu Leu His Leu 40  45  INFORMATION FOR SEQ ID R (i) SEQUENCE CHARACTER (A) LENGTH: 370 (B) TYPE: NUCLEI	(i) SEQUENCE CHARACTERISTI  (A) LENGTH: 299 base  (B) TYPE: NUCLEIC AC  (C) STRANDEDNESS: DC  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE:  (A) ORGANISM: Homo S  (F) TISSUE TYPE: Bra  (ix) FEATURE:  (A) NAME/KEY: Sig_pe  (B) LOCATION: 9014  (C) IDENTIFICATION M  (D) OTHER INFORMATION  (Xi) SEQUENCE DESCRIPTION:  AGCGGA SSTAGCACAG TCCGCCGAGG  GTGGGA GGCTTGAAAC TGTTACAAT  CTG GTT CTG CAA AGT ATG GCT Leu Val Leu Gln Ser Met Ala  -5  ATT GCT GAC TTG TCA GTG AAT  Ile Ala Asp Leu Ser Val Asn  10  AGC TGG AGA AGG GAA GGA GCC Ser Trp Arg Arg Glu Gly Ala  25  GCC TTT CTC CTC CAT TTA CCC  Ala Phe Leu Leu His Leu Pro  40  45  INFORMATION FOR SEQ ID NO:  (i) SEQUENCE CHARACTERIST:  (A) LENGTH: 370 base  (B) TYPE: NUCLEIC AC	(B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapie (F) TISSUE TYPE: Brain  (ix) FEATURE: (A) NAME/KEY: sig_peptic (B) LOCATION: 90143 (C) IDENTIFICATION METHO (D) OTHER INFORMATION:  (xi) SEQUENCE DESCRIPTION: SECUNDARY AND SECUND	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 299 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE:  (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain  (ix) FEATURE:  (A) NAME/KEY: sig_peptide (B) LOCATION: 90143 (C) IDENTIFICATION METHOD: V (D) OTHER INFORMATION: SCORE  (XI) SEQUENCE DESCRIPTION: SEQ ID  AGCGGA SSTAGCACAG TCCGCCGAGC ACAAGCT  METALA  AGCT GAT CTG CAA AGT ATG GCT ACA GGG Leu Val Leu Gln Ser Met Ala Thr Gly  -5  ATT GCT GAC TTG TCA GTG AAT ATG TAT ILE Ala ASP Leu Ser Val ASN Met Tyr 10  15  AGC TGG AGA AGG GAA GGA GCC AGC AGA Ser Trp Arg Arg Glu Gly Ala Ser Arg 25  30  GCC TTT CTC CTC CAT TTA CCC CTT ACA Ala Phe Leu Leu His Leu Pro Leu Thr 40  45  INFORMATION FOR SEQ ID NO: 58:  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 370 base pairs	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 299 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain  (ix) FEATURE: (A) NAME/KEY: sig_peptide (B) LOCATION: 90143 (C) IDENTIFICATION METHOD: Von H (D) OTHER INFORMATION: SCORE 7. Seq LFSL  (xi) SEQUENCE DESCRIPTION: SEQ ID NO:  AGCGGA SSTAGCACAG TCCGCCGAGC ACAAGCTCCA  GTGGGA GGCTTGAAAC TGTTACAAT ATG GCT TTC Met Ala Phe  CTG GTT CTG CAA AGT ATG GCT ACA GGG GCC Leu Val Leu Gln Ser Met Ala Thr Gly Ala -5  ATT GCT GAC TTG TCA GTG AAT ATG TAT AAT ILE Ala Asp Leu Ser Val Asn Met Tyr Asn 10  AGC TGG AGA AGG GAA GGA GCC AGC AGA CAA Ser Trp Arg Arg Glu Gly Ala Ser Arg Gln 25  GCC TTT CTC CTC CAT TTA CCC CTT ACA CAC Ala Phe Leu Leu His Leu Pro Leu Thr His 40  45  INFORMATION FOR SEQ ID NO: 58:  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 370 base pairs (B) TYPE: NUCLEIC ACID	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 299 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE:  (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain  (ix) FEATURE:  (A) NAME/KEY: sig_peptide (B) LOCATION: 90143 (C) IDENTIFICATION METHOD: Von Heijn (D) OTHER INFORMATION: score 7.9  seq LFSLLVLQ  (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 57:  AGCGGA SSTAGCACAG TCCGCCGAGC ACAAGCTCCA GCAT  Met Ala Phe Leu  -15  CTG GTT CTG CAA AGT ATG GCT ACA GGG GCC ACT Leu Val Leu Gln Ser Met Ala Thr Gly Ala Thr -5  ATT GCT GAC TTG TCA GTG AAT ATG TAT AAT CGT Ile Ala Asp Leu Ser Val Asn Met Tyr Asn Arg 10  AGC TGG AGA AGG GAA GGA GCC AGC AGA CAA ATT Ser Trp Arg Arg Glu Gly Ala Ser Arg Gln Ile 25  30  GCC TTT CTC CTC CAT TTA CCC CTT ACA CAC ACA Ala Phe Leu Leu His Leu Pro Leu Thr His Thr 40  45  INFORMATION FOR SEQ ID NO: 58:  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 370 base pairs (B) TYPE: NUCLEIC ACID	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 299 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain  (ix) FEATURE: (A) NAME/KEY: sig_peptide (3) LOCATION: 90143 (C) IDENTIFICATION METHOD: Von Heijne ma (D) OTHER INFORMATION: score 7.9 seq LFSLLVLQSMAT  (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 57:  AGCGGA SSTAGCACAG TCCGCCGAGC ACAAGCTCCA GCATCCCC GTGGGA GGCTTGAAAC TGTTACAAT ATG GCT TTC CTT GGA Met Ala Phe Leu Gly -15  CTG GTT CTG CAA AGT ATG GCT ACA GGG GCC ACT TTC Leu Val Leu Gln Ser Met Ala Thr Gly Ala Thr Phe -5 1  ATT GCT GAC TTG TCA GTG AAT ATG TAT AAT CGT CTT Ile Ala Asp Leu Ser Val Asn Met Tyr Asn Arg Leu 10 15  AGC TGG AGA AGG GAA GGA GCC AGC AGA CAA ATT GCT SET TTP Arg Arg Glu Gly Ala Ser Arg Gln Ile Ala 25 30 35  GCC TTT CTC CTC CAT TTA CCC CTT ACA CAC ACA CAC Ala Phe Leu Leu His Leu Pro Leu Thr His Thr His 40 45 50  INFORMATION FOR SEQ ID NO: 58:  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 370 base pairs (B) TYPE: NUCLEIC ACID	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 299 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain  (ix) FEATURE: (A) NAME/KEY: sig_peptide (B) LOCATION: 90143 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 7.9 seq LFSLLVLQSMATG/AT  (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 57:  AGCGGA SSTAGCACAG TCCGCCGAGC ACAAGCTCCA GCATCCCGTC ACAGCGGA GCCTTGAAAC TGTTACAAT ATG GCT TTC CTT GGA CTC Met Ala Phe Leu Gly Leu -15  CTG GTT CTG CAA AGT ATG GCT ACA GGG GCC ACT TTC CCT Leu Val Leu Gln Ser Met Ala Thr Gly Ala Thr Phe Pro -5  ATT GCT GAC TTG TCA GTG AAT ATG TAT AAT CGT CTT AGA Ile Ala Asp Leu Ser Val Asn Met Tyr Asn Arg Leu Arg 10  AGC TGG AGA AGG GAA GGA GCC AGC AGA CAA ATT GCT TCG Ser TTP Arg Arg Glu Gly Ala Ser Arg Gln Ile Ala Ser 25  GCC TTT CTC CTC CAT TTA CCC CTT ACA CAC ACA CAC GGG Ala Phe Leu Leu His Leu Pro Leu Thr His Thr His Gly 40  45  INFORMATION FOR SEQ ID NO: 58:  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 370 base pairs (9) TYPE: NUCLEIC ACID	(i) SEQUENCE CHARACTERISTICS:    (A) LENGTH: 299 base pairs    (B) TYPE: NUCLEIC ACID    (C) STRANDEDNESS: DOUBLE    (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE:    (A) ORGANISM: Homo Sapiens    (F) TISSUE TYPE: Brain  (ix) FEATURE:    (A) NAME/KEY: sig_peptide    (B) LOCATION: 90143    (C) IDENTIFICATION METHOD: Von Heijne matrix    (D) OTHER INFORMATION: score 7.9	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 299 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain  (ix) FEATURE: (A) NAME/KEY: sig_peptide (B) LOCATION: 90143 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: SCORE 7.9 SEQ LEFSLLVLQSMATG/AT  (XI) SEQUENCE DESCRIPTION: SEQ ID NO: 57:  AGCGGA SSTAGCACAG TCCGCCGAGC ACAAGCTCCA GCATCCCGTC AGGGTTGCAG STGGGA GGCTTGAAAC TGTTACAAT ATG GCT TTC CTT GGA CTC TTC TCT MET ALA PHE LEU Gly LEU PHE SER -15  CTG GTT CTG CAA AGT ATG GCT ACA GGG GCC ACT TTC CCT GAG GAA LEU VAL LEU Gln Ser Met Ala Thr Gly Ala Thr Phe Pro Glu Glu -5 1 5  ATT GCT GAC TTG TCA GTG AAT ATG TAT AAT CGT CTT AGA GCA GTT ILE ALA ASP LEU SER VAL ASN MET TYR ASN ATG LEU ATG ALA VAL 10 15 20  AGC TGG AGA AGG GAA GGA GCC AGC AGA CAA ATT GCT TCG TGT CTG SET TTP ATG ATG GLY ALA SER ATG GLN ILE ALA SER CYS LEU 25 30 35  GCC TTT CTC CTC CAT TTA CCC CTT ACA CAC ACA CAC GGG ALA PHE LEU LEU HIS LEU PRO LEU THR HIS THR HIS GLY 40 45 50  INFORMATION FOR SEQ ID NO: 58:  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 370 base pairs (B) TYPE: NUCLEIC ACID

(D) TOPOLOGY: LINEAR

WO 99/06552		44	<b>.</b>	РСТ/ІВ98/
(ii) MOLEC	CULE TYPE: C	DNA		
(A)	NAL SOURCE: ORGANISM: HO TISSUE TYPE	•		
(B) (C)	NAME/KEY: s: LOCATION: 6:	2226 ION METHOD: V MATION: scor	on Heijne matri e 7.8 ALLVALLFTLIHR/R	
(xi) SEQUE	NCE DESCRIP	TION: SEQ ID	NO: 58:	
AAGTGAGAAA GGAGC	CTTACC AAAGG	CAGTG TACGAAG	AAG GTTCCTGGGA	GACTGTCAGA 60
A ATG AGT TTT TO Met Ser Phe Se -55			G GCG AAC ACA A o Ala Asn Thr T -45	
CCT GTC ACA GGT Pro Val Thr Gly				
GCG GCG GGC ATA Ala Ala Gly Ile -20				Ala Leu
CTA TTT ACT TTG Leu Phe Thr Leu -5				
GAA AGT GAC AGA Glu Ser Asp Arg 10				
ATA TCT GAG AAT Ile Ser Glu Asn	Pro Arg Arg		His Glu Lys Asn	Thr Met
GGA GCA CAA GAG Gly Ala Gln Glu 45				370
	FOR SEQ ID  NCE CHARACTE LENGTH: 336	RISTICS:		
	TYPE: NUCLE			

# (2)

- (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain

,	WUY	9/003:	52						4	5					101	71070/0
		ix) 1	(A) (B) (C) (D)	NAME LOCA IDEN OTHE	ATION NTIFI CR IN	N: 91 CATI	ig_pe i33 (ON M MATIC	BO METHO DN:	DD: V scoi	ce 7. LVLE	7 LSL					
TAT	rcct'	rgg i	AGTT(	CCAC	GA C	rgaa'	)AAT1	G AC	rgtt(	GTGG	GRD	CCAT	AAT 1	TTTC	AAATAC	60
TTG	CCT	ATA '	rtcgʻ	rgtt(	GA GO	GGTT(	CACA		t Se					u Ala	A CTT a Leu	114
							GAC Asp -65									162
							ACA Thr									210
							TCG Ser									258
GGT Gly	GGT Gly	TCC Ser	ATT Ile	TTA Leu -20	CAG Gln	CAC His	TCA Ser	GTG Val	GAT Asp -15	CCC Pro	CTT Leu	GTT Val	TTG Leu	TTC Phe -10	CTA Leu	306
							CCC Pro		-							336
(2)	INF	ORMA'	rion	FOR	SEQ	ID I	NO:	60:								
	(:	i) SI	(A) (B) (C)	LENG TYPE STRA	STH: E: NU ANDEI	394 JCLEI ONES	RIST: base IC AC S: DC	e pa: CID DUBLI								
	(:	ii) I	MOLE	CULE	TYPI	E: CI	DNA									
	(	vi) (	(A)	ORG		4: H	omo S		ens							

- (F) TISSUE TYPE: Brain
- (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
  (B) LOCATION: 266..322
  (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 7.7
  - seq LQLLCCIFTLVLQ/HY
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 60:

ATACTA TAAG	GCTAGAATTG ACT	TGGAGTC AAGGCG	AGTT CCATAGGTTT	AATGCGTGGC 60
ACCTATCTCA	ACAACTGCAA AAT	CACTGTA ACTTGT	AAAA AATCCCAGGG	CCACTCATGG 120
ACAGCCACAT	CTTTAAACCC ACA	CAAAAAT ACAGAC	ACTT TGATTACATG	AAAGTTAGAA 180
GCTCTAAACT	AGAGCAAGGT CAG	TTGTGGT AAGAAG	GTAT TAGATTCAAG	CATCCACAAA 240
AGTTTATAGT	CTGTCAGTCA TAA		GGG ATT TCT TT Gly Ile Ser Le -15	
	T ATT TTC ACT T S Ile Phe Thr L -5			
	T ATT ACC TGC A T Ile Thr Cys I 10		Leu Leu Leu As	
CAG CAG Gln Gln				394

## (2) INFORMATION FOR SEQ ID NO: 61:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 429 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 208..264
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 7.6

seq LLNLLLLSLFAGL/DP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 61:

GAGAATGCCT GCNGAATGAT CGCCCCCCAG GGCGGCTGCC GCCGCTGCCG CTGCTGCTGT 60

TATTGCTACT GCTGCTGCCG CCGCCTCTGC TTCCACTCGG CTCTGACTGG CAGGCARAAA 120

RTGCAACTTG AMSGARGGRH ARGTCTCTGG CAGTGAGTGG AGAGCCTACA TAAAAGAGAG 180

TAAAAGAGGGG CAAAAACCCA GATCAGA ATG CAG GCG ACG TCC AAC CTT CTC AAC 234

Met Gln Ala Thr Ser Asn Leu Leu Asn
-15

					•	•				
								AAG Lys		282
							 	GAT Asp 20	 	330
								CAT His	 	378
								AAG Lys		426
TTG Leu										429

#### (2) INFORMATION FOR SEQ ID NO: 62:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 189 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA-
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 88..180
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 7.6

seq VTLLCGWPGSHWC/AP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 62:

AACAGGCGTA ASKACATGGC CCAGCTCGAT CCCTCCCTTT TGTTCAACAA ACTAAATTCG 60

AGCAGGAGGC TCTAGGATTC CACAGGC ATG ATG AAA TGG AAG CCG GAG GAT CTG 114

Met Met Lys Trp Lys Pro Glu Asp Leu

-30 -25

GGA TCG GTT CCT TGT GAG GCT TTC TCT GTT ACT CTG CTG TGC GGC TGG
Gly Ser Val Pro Cys Glu Aia Phe Ser Val Thr Leu Leu Cys Gly Trp
-20
-15
-10

CCA GGG TCG CAT TGG TGT GCC CCA CCA
Pro Gly Ser His Trp Cys Ala Pro Pro
-5
1

WO 99/06552 48

								7	U						
INFO	ORMA:	rion	FOR	SEQ	ID I	NO:	63:								
<b>(</b> )	L) SE	(A) (B) (C)	LENC TYPE STRA	STH: E: NO ANDEO	243 JCLEI ONESS	base C AC S: DC	e pai CID DUBLE								
( i	.i) N	40LEC	CULE	TYP	E: CI	ANC									
(1)	7i) (	(A)	ORGA	NISN	1: Ho			ens							
(i	.x) E	(A) (B) (C)	NAME LOCA IDEN	ATION NTIFI	N: 10	066 000	5 1etho	D: V	e 7.	6					
к)	:i) S	EQUE	ENCE	DESC	CRIP	: NOI	: SE(	) ID	NO:	63:					
ATCAC					nr Se	er As				sn Le	eu Le				
TTT Phe	GCC Ala	GGA Gly	TTA Leu	GAT Asp 1	CCT Pro	TCC Ser	AAG Lys	ACT Thr 5	CAG Gln	ATT Ile	AGT Ser	CCT Pro	AAA Lys 10	GAA Glu	99
TGG Trp	CAG Gln	GTG Val 15	TAC Tyr	AGC Ser	TCA Ser	GCT Ala	CAG Gln 20	GAT Asp	CCT Pro	GAT Asp	GGG Gly	CGG Arg 25	TGC Cys	ATT Ile	147
ACA Thr	GTT Val 30	GTW Val	GCT Ala	CCA Pro	GAA Glu	CAA Gln 35	AAC Asn	CTG Leu	TGT Cys	TCC Ser	CGG Arg 40	GAT Asp	GCC Ala	AAA Lys	195
															243
		EQUE	NCE (	CHAR	ACTE!	RIST	ICS:	irs							
	(i) (i) (i) (i) (ii) (iii) (iii) (iiii) (iiii) (iiii) (iiiii) (iiiii) (iiiii) (iiiii) (iiiii) (iiiiii) (iiiiii) (iiiiiii) (iiiiiii) (iiiiiiii	(ii) N (iii) N (vi) ( (ix) E (xi) S ATCAGA A: Mc TTT GCC Phe Ala TGG CAG Trp Gln ACA GTT Thr Val 30 AGG CAA Arg Gln 45	(i) SEQUENT (A) (B) (C) (D) (ii) MOLECT (A) (F) (ix) FEATO (A) (B) (C) (D)  (xi) SEQUENT (A)  ATCAGA ATG CA Met G:  TTT GCC GGA Phe Ala Gly  TGG CAG GTG Trp Gln Val 15  ACA GTT GTW Thr Val Val 30  AGG CAA CTT Arg Gln Leu 45  INFORMATION (i) SEQUENT	(i) SEQUENCE ( (A) LENG (B) TYPE (C) STRA (D) TOPO (ii) MOLECULE (vi) ORIGINAL (A) ORGA (F) TISS (ix) FEATURE: (A) NAME (B) LOCA (C) IDEN (D) OTHE  (xi) SEQUENCE  ATCAGA ATG CAG GO Met Gln AI  TTT GCC GGA TTA Phe Ala Gly Leu  TGG CAG GTG TAC Trp Gln Val Tyr 15  ACA GTT GTW GCT Thr Val Val Ala 30  AGG CAA CTT CGC Arg Gln Leu Arg 45  INFORMATION FOR (i) SEQUENCE (	(i) SEQUENCE CHARM  (A) LENGTH:  (B) TYPE: NO  (C) STRANDED  (D) TOPOLOGY  (ii) MOLECULE TYPE  (vi) ORIGINAL SOUND  (A) ORGANISM  (F) TISSUE TO  (ix) FEATURE:  (A) NAME/KEY  (B) LOCATION  (C) IDENTIFE  (D) OTHER IN  (xi) SEQUENCE DESCRATAGA ATG CAG GCG ACC ACC ACC ACC ACC ACC ACC AC	(i) SEQUENCE CHARACTER  (A) BENGTH: 243  (B) TYPE: NUCLER  (C) STRANDEDNESS  (D) TOPOLOGY: LI  (ii) MOLECULE TYPE: CI  (vi) ORIGINAL SOURCE:  (A) ORGANISM: Ho  (F) TISSUE TYPE:  (ix) FEATURE:  (A) NAME/KEY: Si  (B) LOCATION: 10  (C) IDENTIFICATI  (D) OTHER INFORM  (xi) SEQUENCE DESCRIPT  ATCAGA ATG CAG GCG ACG TO  Met Gln Ala Thr Se  TTT GCC GGA TTA GAT CCT  Phe Ala Gly Leu Asp Pro  1  TGG CAG GTG TAC AGC TCA  Trp Gln Val Tyr Ser Ser  15  ACA GTT GTW GCT CCA GAA  Thr Val Val Ala Pro Glu  30  AGG CAA CTT CGC CAA CTA  Arg Gln Leu Arg Gln Leu  45  INFORMATION FOR SEQ ID II  (i) SEQUENCE CHARACTER	(i) SEQUENCE CHARACTERIST:  (A) MENGTH: 243 base  (B) TYPE: NUCLEIC AG  (C) STRANDEDNESS: DG  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE:  (A) ORGANISM: Homo S  (F) TISSUE TYPE: Bree  (ix) FEATURE:  (A) NAME/KEY: sig_pe  (B) LOCATION: 1066  (C) IDENTIFICATION M  (D) OTHER INFORMATION  (xi) SEQUENCE DESCRIPTION:  ATCAGA ATG CAG GCG ACG TCC AF  Met Gln Ala Thr Ser As  —15  TTT GCC GGA TTA GAT CCT TCC  Phe Ala Gly Leu Asp Pro Ser  1  TGG CAG GTG TAC AGC TCA GCT  Trp Gln Val Tyr Ser Ser Ala  15  ACA GTT GTW GCT CCA GAA CAA  Thr Val Val Ala Pro Glu Gln  30  35  AGG CAA CTT CGC CAA CTA CTG  Arg Gln Leu Arg Gln Leu Leu  45  INFORMATION FOR SEQ ID NO:  (i) SEQUENCE CHARACTERIST:	(B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapic (F) TISSUE TYPE: Brain  (ix) FEATURE: (A) NAME/KEY: sig_peptic (B) LOCATION: 1066 (C) IDENTIFICATION METHO (D) OTHER INFORMATION:  (xi) SEQUENCE DESCRIPTION: SEC  ATCAGA ATG CAG GCG ACG TCC AAC COMet Gln Ala Thr Ser Asn Legarian  TTT GCC GGA TTA GAT CCT TCC AAG Phe Ala Gly Leu Asp Pro Ser Lys 1  TGG CAG GTG TAC AGC TCA GCT CAG Trp Gln Val Tyr Ser Ser Ala Gln 15 20  ACA GTT GTW GCT CCA GAA CAA AAC Thr Val Val Ala Pro Glu Gln Asn 30 35  AGG CAA CTT CGC CAA CTA CTG GAA Arg Gln Leu Arg Gln Leu Leu Glu 45 50  INFORMATION FOR SEQ ID NO: 64:  (i) SEQUENCE CHARACTERISTICS:	(i) SEQUENCE CHARACTERISTICS:  (A) MENGTH: 243 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain  (ix) FEATURE: (A) NAME/KEY: sig_peptide (B) LOCATION: 1066 (C) IDENTIFICATION METHOD: V (D) OTHER INFORMATION: SCORE (xi) SEQUENCE DESCRIPTION: SEQ ID  ATCAGA ATG CAG GCG ACG TCC AAC CTT COMET GOTH CO	(i) SEQUENCE CHARACTERISTICS:  (A) BENGTH: 243 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEONESS: DOUBLE  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE:  (A) ORGANISM: Homo Sapiens  (F) TISSUE TYPE: Brain  (ix) FEATURE:  (A) NAME/KEY: sig_peptide  (B) LOCATION: 1066  (C) IDENTIFICATION METHOD: Von F  (D) OTHER INFORMATION: score 7. seq LLNI  (xi) SEQUENCE DESCRIPTION: SEQ ID NO:  ATCAGA ATG CAG GCG ACG TCC AAC CTT CTC AVA  Met Gln Ala Thr Ser Asn Leu Leu As  —15  TTT GCC GGA TTA GAT CCT TCC AAG ACT CAG Phe Ala Gly Leu Asp Pro Ser Lys Thr Gln  1 5  TGG CAG GTG TAC AGC TCA GCT CAG GAT CCT Trp Gln Val Tyr Ser Ser Ala Gln Asp Pro  15 20  ACA GTT GTW GCT CCA GAA CAA AAC CTG TGT Thr Val Val Ala Pro Glu Gln Asn Leu Cys 30 35  AGG CAA CTT CGC CAA CTA CTG GAA AAG GTT Arg Gln Leu Arg Gln Leu Leu Glu Lys Val  45 50  INFORMATION FOR SEQ ID NO: 64:  (i) SEQUENCE CHARACTERISTICS:	(i) SEQUENCE CHARACTERISTICS:	(i) SEQUENCE CHARACTERISTICS:  (A) MENGTH: 243 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE:  (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain  (ix) FEATURE:  (A) NAME/KEY: sig_peptide (B) LOCATION: 1066  (C) IDENTIFICATION METHOD: Von Heijne ma (D) OTHER INFORMATION: score 7.6  seq LLNLLLLSLFAC  (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 63:  ATCAGA ATG CAG GCG ACG TCC AAC CTT CTC AAC CTC CT Met Gln Ala Thr Ser Asn Leu Leu Asn Leu Le  -15  TTT GCC GGA TTA GAT CCT TCC AAG ACT CAG ATT AGT Phe Ala Gly Leu Asp Pro Ser Lys Thr Gln Ile Ser  1  TGG CAG GTG TAC AGC TCA GCT CAG GAT CCT GAT GGG Trp Gln Val Tyr Ser Ser Ala Gln Asp Pro Asp Gly 15  ACA GTT GTW GCT CCA GAA CAA AAC CTG TGT TCC CGG Thr Val Val Ala Pro Glu Gln Asn Leu Cys Ser Arg 30  AGG CAA CTT CGC CAA CTA CTG GAA AAG GTT CAG AAC Arg Gln Leu Arg Gln Leu Leu Glu Lys Val Gln Asn 45  INFORMATION FOR SEQ ID NO: 64:  (i) SEQUENCE CHARACTERISTICS:	(i) SEQUENCE CHARACTERISTICS:  (A) MENGTH: 243 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE:  (A) ORGANISM: Homo Sapiens  (F) TISSUE TYPE: Brain  (ix) FEATURE:  (A) NAME/KEY: sig_peptide  (B) LOCATION: 1066  (C) IDENTIFICATION METHOD: Von Heijne matrix  (D) OTHER INFORMATION: score 7.6  seq_LLNLLLLSLFAGL/DE  (xi) SEQUENCE DESCRIPTION: SEQ_ID_NO: 63:  ATCAGA ATG_CAG_GCG_ACG_TCC_AAC_CTT_CTC_AAC_CTC_CTG_CT  Met_Gln_Ala_Thr_Ser_Asn_Leu_Leu_Asn_Leu_Leu_Leu_Leu_Leu_Leu_Leu_Leu_Leu_Leu	(i) SEQUENCE CHARACTERISTICS:  (A) BENGTH: 243 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE:  (A) ORGANISM: Homo Sapiens  (F) TISSUE TYPE: Brain  (ix) FEATURE:  (A) NAME/KEY: sig_peptide  (B) LOCATION: 1066  (C) IDENTIFICATION METHOD: Von Heijne matrix  (D) OTHER INFORMATION: score 7.6  seq LLNLLLSLFAGL/DP  (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 63:  ATCAGA ATG CAG GCG ACG TCC AAC CTT CTC AAC CTC CTG CTG CTG Met Gln Ala Thr Ser Asn Leu Leu Asn Leu	(i) SEQUENCE CHARACTERISTICS:  (A) MEMGTH: 243 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE:  (A) ORGANISM: Homo Sapiens  (F) TISSUE TYPE: Brain  (ix) FEATURE:  (A) NAME/KEY: sig_peptide  (B) LOCATION: 1066  (C) IDENTIFICATION METHOD: Von Heijne matrix  (D) OTHER INFORMATION: score 7.6  seq LLNLLLLSLFAGL/DP  (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 63:  ATCAGA ATG CAG GCG ACG TCC AAC CTT CTC AAC CTC CTG CTG CTG Met Gln Ala Thr Ser Asn Leu Leu Asn Leu Leu Leu Leu Leu Se:  -15  -10  TTT GCC GGA TTA GAT CCT TCC AAG ACT CAG ATT AGT CCT AAA GAA Phe Ala Gly Leu Asp Pro Ser Lys Thr Gln Ile Ser Pro Lys Glu  1  TGG CAG GTG TAC AGC TCA GCT CAG GAT CCT GAT GGG CGG TGC ATT Trp Gln Val Tyr Ser Ser Ala Gln Asp Pro Asp Gly Arg Cys Ile  15  20  25  ACA GTT GTW GCT CCA GAA CAA AAC CTG TGT TCC CGG GAT GCC AAA Thr Val Val Ala Pro Glu Gln Asn Leu Cys Ser Arg Asp Ala Lys 30  35  40  AGG CAA CTT CGC CAA CTA CTG GAA AAG GTT CAG AAC ATG TCC CGG Arg Gln Leu Arg Gln Leu Leu Glu Lys Val Gln Asn Met Ser Arg 45  INFORMATION FOR SEQ ID NO: 64:  (i) SEQUENCE CHARACTERISTICS:

- (B) TYPE: NUCLEIC ACID
  (C) STRANDEDNESS: DOUBLE
  (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain

(	(B) (C)	URE: NAME/KE LOCATION IDENTIF: OTHER IN	N: 158 ICATION I	301 METHO	DD: \ scoi	on FVII	. 5					
(	xi) SEQU	ENCE DES	CRIPTION	: SE(	Q ID	NO:	64:					
ACAAACA	GAC GMTA	CCATCG C	TTCAGCAG	C AT	CCTC	rcag	ACA	AGAG	CCA (	CTAT	TTCTGA	60
TTCAGAT	CAC CTGT	CATCGA A	GTTTAAAG	A AGO	GGGA	AACA	GGA	GACA	GAA A	ATAC	ACTGAA	120
CCAAAAA	GAT TCAA	LAAGAGC A	AGTGG <b>A</b> AT	C TC	TAAG					r His	C TGG S Trp	175
AAT GAA Asn Glu	ACC ACT Thr Thr -40	ACC TCT Thr Ser	GTT TAT Val Tyr -35	Gln	TAC Tyr	CTT Leu	GGT Gly	TTT Phe -30	CAA Gln	GTT Val	CAA Gln	223
AAA ATT Lys Ile -25	Tyr Pro	TTC CAT Phe His	GAC AAC Asp Asn -20	TGG Trp	AAC Asn	ACT Thr	GCC Ala -15	TGC Cys	TTT Phe	GTC Val	ATC Ile	271
		ATA TTT Ile Phe										319
		CTT GAC Leu Asp										367
		AGT GAA Ser Glu		Pro								397
(2) INF	ORMATION	FOR SEQ	ID NO:	65:								
(	(A) (B) (C)	NCE CHAR LENGTH: TYPE: NO STRANDE TOPOLOG	182 base UCLEIC A DNESS: D	e pai CID OUBLE								
(	ii) MOLE	CULE TYP	E: CDNA									
(	(A)	INAL SOU ORGANIS TISSUE	H: Homo	-	ens							
(	(B) (C)	URE: NAME/KE LOCATIO IDENTIF OTHER I	N: 781 ICATION	76 METHO	OD: 1		. 4					

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 65:

ATCGACTGTG AGCTGCGGCA GAGAGCAGAG GCGGCGGCGC GGGACCTGCA GTCGCCAGGG	60
ATTCCCTCCA GGTGACG ATG CTC TGG TTC TCC GGC GTC GGG GCT CTG GCT  Met Leu Trp Phe Ser Gly Val Gly Ala Leu Ala  -30  -25	110
GAG CGT TAC TGC CGC CGC TCG CCT GGG ATT ACG TGC TGC GTC TTG CTG Glu Arg Tyr Cys Arg Arg Ser Pro Gly Ile Thr Cys Cys Val Leu Leu -20 -15 -10	158
CTA CTC AAT TGC TCA GGG GTC TGG Leu Leu Asn Cys Ser Gly Val Trp -5	182
(2) INFORMATION FOR SEQ ID NO: 66:  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 256 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain  (ix) FEATURE: (A) NAME/KEY: sig_peptide (B) LOCATION: 164238 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 7.3 seq LIFFLNVTQLVRG/RG  (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 66:	
AGTAAATAAA AAGTTTGCTT TATTAAATTA TGTTTAGATA GTGSTTATAG TGCTTTACCC	60
CTTCAAAATA GTAACTTCTA TCAATCATTT AGGATGTGTG TCAGACTATT CTGTGTCCTT	120
TAAGTGTGTK AACTAGTTTT AACCCTCTGC AAATATCTGA GGT ATG CTC TTT TTA Met Leu Phe Leu -25	175
CAG ATG GGA AAA CAA TCT TGG ACT TTA ATA TTT TTT CTT AAT GTT ACA Gln Met Gly Lys Gln Ser Trp Thr Leu Ile Phe Phe Leu Asn Val Thr -20 -15 -10	223
CAA TTA GTA AGA GGC AGG GGG CCA GGC GGA CGG Gln Leu Val Arg Gly Arg Gly Pro Gly Gly Arg -5 1 5	256

(2)	INFORM	MATION	FOR	SEQ	ID !	NO: (	67 :						
	(i)	(B) (C)	NCE ( LENC TYPE STRA TOPO	STH: E: NU ANDEI	126 JCLEI ONESS	base C AC S: DC	e pai CID OUBLE						
	(ii)	MOLE	CULE	TYPE	E: CI	ANC							
	(vi)		INAL ORGA TISS	NISM	1: Hc		-	ens					
	(ix)	(B) (C)	URE: NAME LOCA IDEN OTHE	TION TIFI	: 19 :CATI	099 ON M	) 1ETHC	D: \ scoi	e 7.	2	itrix SL/AS		
	(xi)	SEQU	ENCE	DESC	CRIPT	:NOI	SEQ	Q ID	NO:	67:			ŕ
AAT	rgattac	GAGA	TTAT								GGG Gly		51
	GTG CA Val Gl -15												99
	TCC TI Ser Ph												126
(2)	INFORM	MATION SEQUE											
	(1)	(A) (B) (C)	LENC TYPE STRA	STH: E: NU ANDEI	117 JCLEI NESS	base IC AC S: DC	e pai CID OUBLE						
	(ii)	MOLE	CULE	TYPE	E: CI	ANC							
	(vi)		INAL ORGA TISS	RIMA	1: Ho		-	ens					
	(ix)	(B) (C)	URE: NAME LOCA IDEN	ATION ATIF:	1: 46 [CAT]	88 1 NO	7 METH(	DD: 1	re 7		atri: FG/I!		
	(xi)	SEQU	ZNCE	DESC	CRIP'	TION	: SE	Q ID	NO:	68:			

AGCA	10000	,11 1	GGGF	NONOF	M AC	CIAL	11600	. 1 ^\	AGCC 1	CAT	CCT.				p Se	
												CAC His				105
		GAA Glu														117
(2)	INFO	RMAT	CION	FOR	SEQ	ID N	10: 6	59:								
	(i	.) SE	(A) (B) (C)	LENG TYPE STRA	HARA TH: : NU :NDED LOGY	445 CLEI NESS	base C AC : DC	pai ID UBLE								
	(i	.i) M	OLEC	ULE	TYPE	: CE	ANG									
	( v	i) C	(A)	ORGA	SOUR NISM UE T	: Ho			ens							
	i)	.x) F	(B) (C)	NAME LOCA IDEN	KEY TION TIFI R IN	: 11 CATI	0.2	:65 IETHO )N:	D: V	e 7	_	ne ma CYFAE				
	()	i) S	EQUE	NCE	DESC	RIPT	: NOI	SEC	OI O	NO:	69:					
ATGO	CACC	ATG P	TAT	TTTT	AT AC	CACGI	TGT	S TTA	AACT <i>I</i>	ACTG	TAA	ACAC <i>i</i>	ATT (	STCT	CTTT	A 60
TAT	TCT	TTG (	CAGGA	AAGT 1	C AC	AAA	AAAGT	GT(	CACG	TTTT	AAT	CTGC			AC AT. sp Il-	е
AGT Ser	GGA Gly	TTA Leu	ATT Ile	CCT Pro -45	GGT Gly	CTA Leu	GTG Val	TCT Ser	ACA Thr -40	TTC Phe	ATA Ile	CTT Leu	TTG Leu	TCT Ser -35	AKH Xaa	166
												TCT				214
												TAT Tyr -5				262
												GCA Ala				310
A.A.T	TAT	ACC	ACA	TTT	TGG	GGA	GCT	TGC	TTT	GCC	TAT	ATA	GTT	GAT	CAG	358

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Asn Tyr Thr Thr Phe Trp Gly Ala Cys Phe Ala Tyr Ile Val Asp Gln 20 25 30	
TGT AAA GAA CRS DKA CAA AAA ACA ATT CGA ATA GCT ATC ATT GAC TTT Cys Lys Glu Xaa Xaa Gln Lys Thr Ile Arg Ile Ala Ile Ile Asp Phe 35 40 45	406
CTA CTT GGA CTT GTT ACT GGA CTA ACA GTA CTG TCA TCT Leu Leu Gly Leu Val Thr Gly Leu Thr Val Leu Ser Ser 50 55 60	445
(2) INFORMATION FOR SEQ ID NO: 70:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 244 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: CDNA	
<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F; TISSUE TYPE: Brain</pre>	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 137226     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 7</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 70:	
ACTITITGAA TITGTTGCTG GTACAGTTGC ATGTATTCTC TTAAAATTAT TITGAGGCCT	60
CATATCTGGT TATTTCTCCT TTCTCATTCC TTATCTTGCG TGTTTTTACC TTTTTTCAT	120
AACTAAGTTT TTGAGG ATG TWA GTG TTC TTT TCA AAG AAC CGG TTC GAA ATG Met Xaa Val Phe Phe Ser Lys Asn Arg Phe Glu Met -30 -25 -20	172
TAC TTT TCT TTG CTA CTT TTT GTT ATT TTA TTG ATC ACA TCT TTA ATC Tyr Phe Ser Leu Leu Phe Val Ile Leu Leu Ile Thr Ser Leu Ile +15 -10 -5	220
TTT TGT TCT CTA TAC GTG GCG CGT Phe Cys Ser Leu Tyr Val Ala Arg 1 5	244
(2) INFORMATION FOR SEQ ID NO: 71:	
(i) SEQUENCE CHARACTERISTICS:	
(A) LENGTH: 390 base pairs (B) TYPE: NUCLEIC ACID (C: STRANDEDNESS: DOUBLE	

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 289..357
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 6.9

seq SLSLLASHHSVSC/SN

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 71:

AAGTACAAAG CCTACTCCAA AGACTGCAGC TTGAAGATAA AAGAGAGCAC TGCGTCCTTT 60

TGAAAATAAA GGCAGACACA AAGAAGAAAG GAGCTACCTT ACCCCAGCAT ATACCTGCGG 120

GATGTTCTCT CCAGTTCATT TTTACCTGGT GTCTTGAAAT CCGAGCAATT CCTAAAAAGG 180

CATTTTTGCG AGCCCTTGTG GACTATACCA GTGACAGTGC TGAAAAGCGC AGGCTACAGG 240

AGCTGTGCAG TAAACAAGGG GCAGCCGATT ATAGCCGCTT TGTACGAG ATG CCT GTG 297

Met Pro Val

CCT GCT TGT TGG ATC TCC TCC TCG CTT TCC CTT CTT GCC AGC CAC CAC Pro Ala Cys Trp Ile Ser Ser Ser Leu Ser Leu Leu Ala Ser His His -20 -15 -10 -5

TCA GTC TCC TGC TCG AAC ATC TTC CTA AAC TTC AAC CCA GAC CGG 390

Ser Val Ser Cys Ser Asn Ile Phe Leu Asn Phe Asn Pro Asp Arg 10

- (2) INFORMATION FOR SEQ ID NO: 72:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 374 base pairs
    - (E) TYPE: NUCLEIC ACID
    - (C) STRANDEDNESS: DOUBLE
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: CDNA
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: 198..260
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 6.9

seq LLACGSLLPGLWQ/HL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 72:

ATATATTTCT GAGGCAGTAC CCATCTCACT TGTAAACTTA AAAGACACCG CAGAGATTTG	60
AGGGACTCAG AAGTCAAATA GAGTAGGTTA AAAACCTCTT ATTTTTCAAA TTAATTGTTT	120
TAAGAAACAA GCATACCTGT GTAAGTGAAA TATCTTAATT TGTGTTGAAT CAAGTTAGGA	180
GACAGAGATT CTCATGA ATG TGT CCT GTG TTC TCA AAG CAG CTG CTA GCC  Met Cys Pro Val Phe Ser Lys Gln Leu Leu Ala  -20  -15	230
TGT GGG TCT CTC CTA CCT GGG TTA TGG CAG CAC CTC ACA GCC AAT CAC Cys Gly Ser Leu Leu Pro Gly Leu Trp Gln His Leu Thr Ala Asn His -10 -5 1 5	278
TGG CCT CCA TTC TCC SCT TTC CTC TGT ACA GTT TGC TCT GGT TCC TCA Trp Pro Pro Pro Phe Ser Xaa Phe Leu Cys Thr Val Cys Ser Gly Ser Ser 10 15 20	326
GAG CAG ATT TCC GAG TAT ACT GCT TCA GCC ACG CCC CCA CTG TGC CTG Glu Gln Ile Ser Glu Tyr Thr Ala Ser Ala Thr Pro Pro Leu Cys Leu 25 30 35	374
(2) INFORMATION FOR SEQ ID NO: 73:  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 416 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain  (ix) FEATURE: (A) NAME/KEY: sig_peptide (B) LOCATION: 33260 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 6.9 seq LLPLSAWPPWAWH/HH  (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 73:	
ACGCTCAGCA GGTCCACTCC CGTGTTCCGG TC ATG GCT TTA ACA ATT CAT GGG Met Ala Leu Thr Ile His Gly -75 -70	53
GAA AGA ATG CGC CCC GAT TGG GAG AGC CCC TGG ATC ACG TCT TCC CAA Glu Arg Met Arg Pro Asp Trp Glu Ser Pro Trp Ile Thr Ser Ser Gln -65 -60 -55	101
GCT CAG TCC GT% TCT CTT GGA GGG AGT CCG TCC TCG AGG GGC CCT CTG Ala Gln Ser Leu Gly Gly Ser Pro Ser Ser Arg Gly Pro Leu	149

GTG	ccc	AGG	GGA	GAG	TAT	CTT	GCG	TCC	TGT	CCT	GAG	GGC	GTC	CGC	TCA	197
Val	Pro	Arg -35	Gly	Glu	Tyr	Leu	Ala -30	Ser	Cys	Pro	Glu	Gly -25	Val	Arg	Ser	
CAC His	AGC Ser -20	CAC His	CTG Leu	CTC Leu	CCC Pro	CGC Arg -15	TCC Ser	CTC Leu	CTT Leu	CCC Pro	TTG Leu -10	TCA Ser	GCA Ala	TGG Trp	CCA Pro	245
CCG Pro -5	TGG Trp	GCC Ala	TGG Trp	CAT His	CAC His 1	CAT His	GGG Gly	CCT Pro	GGC Gly 5	ACA Thr	CAG <sup>.</sup> Gln	TCC Ser	CTC Leu	GTG Val 10	GGC Gly	293
TGC Cys	CTT Leu	TGT Cys	GCC Ala 15	ATG Met	AGC Ser	CCA Pro	CTG Leu	CTG Leu 20	CCG Pro	ACT Thr	CAC His	CTG Leu	TCC Ser 25	CTC Leu	CCA Pro	341
GTA Val	CTG Leu	GAA Glu 30	CCT Pro	TCT Ser	GGA Gly	ACA Thr	CCA Pro 35	GCA Ala	CTA Leu	AAA Lys	GAT Asp	AGG Arg 40	AGG Arg	CCC Pro	TGT Cys	389
						CCC Pro 50										416
	i) i () () ()	.) SE .i) & .i) C .x) E	CQUEN (A) (B) (C) (D) IOLEC ORIGI (A) (F) FEATU (B) (C) (D)	TYPE STRA TOPO CULE NAL ORGA TISS IRE: NAME LOCA OTHE	CHARACTH: C: NU NDEC LOGY TYPE SOUF NISM GUE T C/KEY ATION HTIFI CR IN	i: Ho YPE: : si i: 11 CATI ICATI	RISTI base C AC S: DC NEAR DNA DNA Bra S Bra CON M MATIC	CS: pai ID UBLE in eptic	ens le DD: V scor seq ) ID	e 6. ILIA NO:	8 ASSLE 74:	PTLSF	IP/AF	<b>,</b>		
AAGA	AGTC#	ACA /	ATG ( det <i>A</i>	Ala A	GCC # Ala # -90	AGA 1 Arg I	TTC # Phe #	AGG 1	Cys (	GGC ( Gly H -85	CAT 1 His I	TTG :	rgt ( Cys \	GTC ( Val I	CCC Pro -80	49
GAG Glu	GTT Val	CCT Pro	CGC Ang	GGG Gly -75	CCG Pro	GCA Ala	TCC Ser	CAC His	GCC Ala -70	GAG Glu	GGT Gly	GGT Gly	GGT Gly	GGC Gly -65	AGG Arg	97
CTT Leu	TCC Ser	AGA Arg	AAG Lya	GCA Ala	GCA Ala	CAC His	CAG Gln	GCT Ala	CAG Gln	CTC Leu	TGC Cys	TGG Trp	CGA Arg	GCA Ala	GGA Gly	145

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WO 99/06552		57		PCT/IB98/0
-60		-55	-50	
GGC GAC GGC AGA Gly Asp Gly Arg -45	GGA AAC TTC AAC Gly Asn Phe Asn -40	Pro Met Asn Pl	TC CTG GTT GCG he Leu Val Ala -35	GGG 193 Gly
	TCC TGC CAC TCA Ser Cys His Ser -25	Pro Pro Leu Le		
CCA AGA ATC CTC Pro Arg Ile Leu -15	ATA GCG TCC TCT Ile Ala Ser Ser -10	CTC CCC ACT CT Leu Pro Thr Le	FC TCC CAT CCC eu Ser His Pro	GCG 289 Ala 1
CCT GGG Pro Gly				295
(2) INFORMATION	FOR SEQ ID NO:	75:		
(A) (B) (C)	NCE CHARACTERIST LENGTH: 361 bas TYPE: NUCLEIC A STRANDEDNESS: DO TOPOLOGY: LINEA	e pairs CID OUBLE		
(ii) MOLE	CULE TYPE: CDNA			
(A)	INAL SOURCE: ORGANISM: Homo TISSUE TYPE: Br			
(B) (C)	URE: NAME/KEY: sig_p LOCATION: 101 IDENTIFICATION OTHER INFORMATION	187 METHOD: Von Hei ON: score 6.8	ijne matrix CSCFYTQP/HP	
(xi) SEQU	ENCE DESCRIPTION	: SEQ ID NO: 7	5:	
ACTCTCCCCT CCCC	TCCCCG GCACTGCAG	C ACCAGCCGTC TO	GCAGCTCCG GCCGC	CCACTT 60
GCGCCTCTCC AGCC	TCCGCA GCCCAACCG		TG GCC AGC ACC et Ala Ser Thr	
	GAG AAG ATG AAG Glu Lys Met Lys -20			
	TAC ACA CAG CCG			
	GTG AAG CAG CTG Val Lys Gln Leu 15	Asp Lys Arg A		

WO 99/06552	50	PCT/IB98/01236

36														
						TCC Ser								307
						AAG Lys								355
CAA Gln														361

## (2) INFORMATION FOR SEQ ID NO: 76:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 361 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA

-30

- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 2..343
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 6.7

seq LIPMAILLGQTQS/NS

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 76:
- A ATG TTA CAA GTC TAT GGA AAG CCA GTT TAT CAG GGC CAT CGA AGC ACT

  Met Leu Gln Val Tyr Gly Lys Pro Val Tyr Gln Gly His Arg Ser Thr

  -110 -105 -100
- CTT AAA AAA GGA CCA TAT CTC AGA TTT AAT TCT CCA TCT CCT AAG TCC 97
  Leu Lys Lys Gly Pro Tyr Leu Arg Phe Asn Ser Pro Ser Pro Lys Ser
- AGA CCA CAG AGA CCA AAA GTA ATA GAA CGA GTT AAA GGC ACT AAG GTA
  Arg Pro Gin Arg Pro Lys Val Ile Glu Arg Val Lys Gly Thr Lys Val
  -80 -75 -70
- AAG TCA ATA AGA ACA CAG ACT GAC TTC TAT GCA ACA AAA CCT AAG AAG
  Lys Ser Ile Arg Thr Gln Thr Asp Phe Tyr Ala Thr Lys Pro Lys Lys
  -65
  -55
- ATG GAT TOT AAA ATG AAA CAT TOT GTT CCT GTG TTA CCT CAT GGC GAT

  Met Asp Ser Lys Met Lys His Ser Val Pro Val Leu Pro His Gly Asp

  -50 -45 -40 -35
- CAG CAA TAT TTG TTC AGC CCA AGT AGA GAA ATG CCT ACT TTT TCA GGT Gln Gln Tyr Leu Phe Ser Pro Ser Arg Glu Met Pro Thr Phe Ser Gly

-25

					CTG Leu											337
					ACC Thr											361
(2)			QUEN (A) (B) (C)	ICE C LENG TYPE STRA	SEQ CHARA TH: : NU .NDED	CTER 388 CLEI NESS	DOSE DO	CS: pai ID UBLE								
	•		RIGI (A)	NAL ORGA	TYPE SOUR NISM UE T	CE:	mo S		ns							
	( <u>i</u>	ж) <b>F</b>	(A) (B) (C)	NAME LOCA IDEN	:/KEY TION TIFI :R IN	: 8. CATI	.361 ON M	ETHO	D: V	e 6.	7		trix SA/RF			
	( x	i) S	EQUE	NCE	DESC	RIPI	'ION:	SEC	) ID	NO:	77:					
AGC#		ATG	TCT	GTA	CTT Leu	GAA Glu	ATC	AGT	GGA	ATG	ATA Ile					49
AAC	AGC	ATG Met	TCT Ser	GTA Val CCA	CTT Leu -115 GGA Gly	GAA Glu i	ATC Ile GGA	AGT Ser	GGA Gly CAG	ATG Met -110	ATA Ile )	Met GGA	Asn AAT	Arg GCA	Val -105 GTC	<b>49</b> 97
AAC Asn TCT	AGAC AGC Ser	ATG Met CAT His	TCT Ser ATA Ile	GTA Val CCA Pro -100	CTT Leu -115 GGA Gly	GAA Glu ATA Ile	ATC Ile GGA Gly	AGT Ser TAC Tyr	GGA Gly CAG Gln -95	ATG Met -110 ATT Ile	ATA Ile TTT Phe	Met GGA Gly CTT	Asn AAT Asn TCT	Arg GCA Ala -90 CAA	Val -105 GTC Val	
AAC Asn TCT Ser	AGAC AGC Ser CTC Leu	ATG Met CAT His ATA Ile	TCT Ser ATA Ile CTG Leu -85	GTA Val CCA Pro -100 GGT Gly	CTT Leu -115 GGA Gly	GAA Glu ATA Ile ACT Thr	ATC Ile GGA Gly CCA Pro	AGT Ser TAC Tyr TTT Phe -80 CAT	GGA Gly CAG Gln -95 GTT Val	ATG Met -110 ATT Ile TTC Phe	ATA Ile TTT Phe CGA Arg	GGA Gly CTT Leu	ASN AAT ASN TCT Ser -75 CTT	GCA Ala -90 CAA Gln	Val -105 GTC Val GCT Ala	97
AAC Asn TCT Ser ACA Thr	AGAC Ser CTC Leu GAC Asp	ATG Met CAT His ATA Ile TTG Leu -70	TCT Ser ATA Ile CTG Leu -85 GAA Glu	GTA Val CCA Pro -100 GGT Gly CAA Gln	CTT Leu -115 GGA Gly ) TTA Leu	GAA Glu ATA Ile ACT Thr	ATC Ile GGA Gly CCA Pro GCA Ala -65 GAT	AGT Ser TAC Tyr TTT Phe -80 CAT His	GGA Gly CAG Gln -95 GTT Val TCT Ser	ATG Met -110 ATT Ile TTC Phe GCT Ala	ATA Ile  TTT Phe  CGA Arg  TCA Ser	GGA Gly CTT Leu GAA Glu -60	ASN  AAT ASN  TCT Ser -75  CTT Leu  ATG	GCA Ala -90 CAA Gln TAT Tyr	Val -105 GTC Val GCT Ala GTG Val	97 145
AAC Asn TCT Ser ACA Thr ATT Ile	AGAC Ser CTC Leu GAC Asp GCA Ala -55	ATG Met CAT His ATA Ile TTG Leu -70 TTT Fhe	TCT Ser ATA Ile CTG Leu -85 GAA Glu GGT Gly	GTA Val CCA Pro -100 GGT Gly CAA Gln TCT Ser	CTT Leu -115 GGA Gly TTA Leu CTC Leu	GAA Glu ATA Ile ACT Thr ACA Thr GAA Glu -50	ATC Ile  GGA Gly  CCA Pro  GCA Ala -65  GAT Asp	AGT Ser TAC Tyr TTT Phe -80 CAT His	GGA Gly CAG Gln -95 GTT Val TCT Ser ATA Ile	ATG Met -110 ATT Ile TTC Phe GCT Ala GTT Val	TTT Phe  CGA Arg  TCA Ser  CTT Leu -45	GGA Gly CTT Leu GAA Glu -60 TCT Ser	ASN AAT ASN TCT Ser -75 CTT Leu ATG Met	GCA Ala -90 CAA Gln TAT Tyr GTT Val	Val -105 GTC Val GCT Ala GTG Val ATA Ile	97 145 193

Leu	Phe	Gly	His	Leu	Thr	Ser	Ala	Arg	Arg	Ala	Arg	Lys 5	Ser	Glu	Val	385
CCT Pro				•												388
(2)	INFO	ORMA?	rion	FOR	SEQ	ID 1	10: 7	78:								
	<b>(</b> )	L) SE	(B) (C)	LENG TYPE STRA	CHARA STH: C: NU NDED	291 CLEI NESS	base C AC : DC	e pai CID OUBLE								
	į )	i) 1	OLEC	CULE	TYPE	: C	NA									
	7)	/i) (		ORGA	SOUF NISM SUE T	I: Ho		-	ens							
	<b>(</b> )	.x) E	(B) (C)	NAME LOCA IDEN	C/KEY TION TIFI CR IN	: 79 CATI	028 ON M	5 SETHO	D: V	e 6.						
	()	(i) S	SEQUE	ENCE	DESC	RIPT	CION:	SEC	Q ID	NO:	78:					
AAG!	ACT (F)	CGA (	CTGG	\AGA(	GG AC	SAAAC	SAACT	r GC	ATGCT	TTGT	AAC	GCC	CGG F	\TGG(	GAGTGT	60
ATTO	CTT	TTT ?	rtga.	AGAT							GCT Ala					111
GAG Glu	AAG Lys	CTG Leu	GTG Val -55	GGA Gly	TAT Tyr	TCT Ser	GCC Ala	GTG Val -50	TAT Tyr	AGA Arg	GTC Val	TGT Cys	TTT Phe -45	GGA Gly	ATG Met	159
			TTC Phe													207
AGC Ser	AAA Lys -25	AGT Ser	TGT Cys	AGA Arg	GCT Ala	CAT His -20	ATT Ile	CAC His	AAT Asn	GGC Gly	TTT Phe -15	TGG Trp	TTC Phe	TTT Phe	AAA Lys	255
			TTG Leu													291

- (2) INFORMATION FOR SEQ ID NO: 79:
  - (1) SEQUENCE CHARACTERISTICS:

WO 99/06552	61	PCT
(B) (C)	LENGTH: 402 base pairs TYPE: NUCLEIC ACID STRANDEDNESS: DOUBLE TOPOLOGY: LINEAR	
(ii) MOLE	CULE TYPE: CDNA	
(A)	INAL SOURCE: ORGANISM: Homo Sapiens TISSUE TYPE: Brain	
(B) (C)	URE:  NAME/KEY: sig_peptide  LOCATION: 61372  IDENTIFICATION METHOD: Von Heijne matrix  OTHER INFORMATION: score 6.6  seq HFSHVVWFHPTWA/QQ	
(xi) SEQU	ENCE DESCRIPTION: SEQ ID NO: 79:	
ATTTTCCCGG GTCT	TCTCCA GCTGCCACCG CTTTACTGCA AAACTGACGG GCGCAA	AAAAC
ATG AGT GAC TCC Met Ser Asp Ser	GCG GGA GGG CGC GCT GGT CTC CGG CGT TAC CCC A Ala Gly Gly Arg Ala Gly Leu Arg Arg Tyr Pro 1 -100 -95 -90	AAG Lys
CTC CCA GTG TGG Leu Pro Val Trp -85	GTG GTG GAG GAT CAT CAG GAG GTT CTA CCC TTT A Val Val Glu Asp His Gln Glu Val Leu Pro Phe : -80 -75	ATA Ile
TAC CGG GCC ATA Tyr Arg Ala Ile -70	GGC TCA AAG CAT CTT CCT GCC AGT AAT GTA AGT CGly Ser Lys His Leu Pro Ala Ser Asn Val Ser 165	ſTT Phe
TTA CAT TTC GAC Leu His Phe Asp	TCA CAT CCA GAC CTC CTT ATT CCT GTG AAT ATG C Ser His Pro Asp Leu Leu Ile Pro Val Asn Met 1	CCA Pro

60

108

156

204

252

300

348

402

77

(2) INFORMATION FOR SEQ ID NO: 80:

CAC TTT

His Phe 10

> (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 338 base pairs

-50

GCA GAC ACC GTG TTT GAT AAG GAA ACA CTC TTT GGA GAA TTA AGT ATT

Ala Asp Thr Val Phe Asp Lys Glu Thr Leu Phe Gly Glu Leu Ser Ile

GAA AAT TGG ATT ATG CCT GCA GTT TAT GCT GGC CAT TTT TCA CAT GTA

Glu Asn Trp Ile Met Pro Ala Val Tyr Ala Gly His Phe Ser His Val

GTA TGG TTT CAT CCC ACA TGG GCT CAG CAG ATC AGA GAG GGC AGA CAC Val Trp Phe His Pro Thr Trp Ala Gln Gln Ile Arg Glu Gly Arg His

-15

(B) TYPE: NUCLEIC ACID(C) STRANDEDNESS: DOUBLE(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens
(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 12..152

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 6.3

seq SSCVLLTALVALA/AY

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 80:

<u>A</u> AA!	ACGC	STG A			Cys			s Vai		CTC Leu	50
				TTT Phe							98
				TCG Ser							146
				GTC Val							194
				CTG Leu 20							242
				CAS Xaa							290
				TTT Phe			Lys	Ala			338

#### (2) INFORMATION FOR SEQ ID NO: 81:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 229 base pairs

(3) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

				ORG!				-	ens							
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 14139     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 6.1</pre>																
	( 2	(i)	SEQU	ENCE	DES	CRIPT	CION:	: SE(	Q ID	NO:	81:					
ATT:	AAAG!	rca .		ATG A	Ile					Ile 1						49
CGA Arg -30	GTC Val	CTG Leu	CTG Leu	TCG Ser	GGA Gly -25	AGG Arg	GAG Glu	ATG Met	TTT Phe	CCT Pro -20	GCT Ala	TCC Ser	GTC Val	CGT Arg	GCT Ala -15	97
CCT Pro	GAC Asp	CTG Leu	GCG	GTG Val -10	GCC Ala	CTG Leu	TCC Ser	CTG Leu	CTA Leu -5	CCT Pro	GCG Ala	TGG Trp	ACA Thr	GAG Glu 1	TCT Ser	145
CCA Pro	ACA Thr	CGC Arg 5	GGC Gly	AGC Ser	CAC His	CAG Gln	AGC Ser 10	CAG Gln	GCC Ala	CGA Arg	GCG Ala	CAC His 15	AGC Ser	CGT Arg	GCA Ala	193
				AGC Ser												229
(2)	INFO	ORMA	TION	FOR	SEQ	ID 1	NO: {	32:								
	i)	L) S	(A) (B) (C)	NCE ( LEN( TYPE STRA TOP(	STH: E: NO ANDEI	249 UCLEI DNESS	base C AC S: DC	pai CID OUBLE								
	( :	i)	MOLE	CULE	TYP	E: CI	ANC									
	(1	vi) (	(A)	INAL ORGA TISS	ANIS	M: Ho			ens							
	(:	ix)	(A) (B) (C)	URE: NAM! LOC! I DE! OTH:	ATION SITE	N: 70 ICAT	022 ION N	28 1ETH(	DD: 7	ce 6	Heijr LLLL					

(mi) SEQUENCE DESCRIPTION: SEQ ID NO: 82:

GGTG	<b>A</b> ATG1				ST AG /s Se -5	r Al				ys I							111
ATT A										Lys					n G		159
CCT G		la '					Lys							u Le			207
TTA G											Va.						249
(2) I	INFOR	TAM	ION	FOR	SEQ	ID N	O: 8	3:									
	(i)	_	-		HARA												
					TH: : NU				Lrs								
					NDED				Ε								
	(ii				TYPE												
	/	٠ .		17 T	COUR	an.											
	(V1	(	(A)	ORGA	SOUR NISM UE T	: Ho		-	ens								
	(ix	(	(A) (B) (C)	NAME LOCA IDEN	/KEY TION TIFI R IN	: 11 CATI	02 ON M	29 ETHO	DD: sco	re 5	. 9						
									seq	VCS.	ALCS	SLGE	VRP/	Œ			
	(xi	.) SI	EQUE	NCE	DESC	RIPT	ION:	SE	O IC	NO:	83	:					
ACCCT	rgcte	G G	CGGG	AAGG	SC GG	CGCC	CCGG	CC	GAGG	TGGC	GG	CGGC	TCCT	CAC	ATG	GGAG	60
AAGAA	AGTTG	T C	CATG	TTC	AC AC	TGGG	TGAA	. GG	AAGC	TGAA	. AC	CACA	1			GAG Glu	118
TCC T Ser S	Ser M												a Il				166
AAG 0 Lys A	GAC C Asp E -20	Pro	CTG Leu	GTG Val	CAG Gln	GAG Glu -15	CAG Gln	GTC Val	TGC Cys	AGT Ser	GC Al	a Le	G TG u Cy	C TO S Se	CC C	TC eu	214
GGG G Gly G -5										, Ala				u Ty			262
cas c	CAS A	ATG .	ACA	AGC	TGG	CAC	ACC	CGG									299

Arg Xaa Met Thr Ser Trp His Thr Arg 15 20

(2) INFORMATION FOR S	SEQ I	טא ט	: 84	:
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- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 252 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 76..204
  - (C) IDENTIFICATION METHOD: Von Heijne matrix

65

(D) OTHER INFORMATION: score 5.9

seq VFLFHCTSGLSSC/KC

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 84:

# AGTARAACAA AAGGATATGC ACACACACAT ATTTAAATAC ATGTAGTTTT TTGCATAAAT 60

- TATCACTGAG AGGAA ATG CAA GAA ACT GAT TGT AAT AAA CGC TGG GGA AGG 111

  Met Gln Glu Thr Asp Cys Asn Lys Arg Trp Gly Arg
  -40 -35
- GGC CTG GGT GGC CTG TGG TCA GAA ACA GGA AGG AGA TTT CAT TGC AAA 159
  Gly Leu Gly Gly Leu Trp Ser Glu Thr Gly Arg Arg Phe His Cys Lys
  -30 -25 -20
- TCT TTT GTA TTT CTT TTT CAC TGT ACT TCT GGA TTA TCT TCA TGC AAA 207 Ser Phe Val Phe Leu Phe His Cys Thr Ser Gly Leu Ser Ser Cys Lys
- TGT TCT AAA AAG CAT TYM AAA TAT TGC TTC TGT TTT GTG GCA AGT

  Cys Ser Lys Lys His Xaa Lys Tyr Cys Phe Cys Phe Val Ala Ser

  5 10 15
- (2) INFORMATION FOR SEQ ID NO: 85:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 366 base pairs
    - (3) TYPE: NUCLEIC ACID
    - (C) STRANDEDNESS: DOUBLE
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: CDNA
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain	(F)	TISSUE	TYPE:	Brain
------------------------	-----	--------	-------	-------

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 232..282
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5.8

seq VPWLSSTVSCAQG/LR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 85:

ATGAGTGGTC MGGAGATAAC ACTTAATGCT TTATTCTTAA GTGTTGGAAG GAGCAAGTAA 60 GTGGTCTAGT GAGCTGTTTT TAGAGGAACT GTATAATATG TAACACATTG TCATTATATT 120 CACTAACTCC CAAAGTATTC TTGAGATATT GANACAAAAC AAAGAGCTTG AATAGAAACC 180 CTGAGCAACA ATGTATTTAC TTTCCACTTG CAGCAGAACT TGGCCTTTCA G ATG CTC 237 Met Leu CTT GAA GTG CCT TGG CTT AGC AGT ACT GTC TCT TGT GCC CAG GGT CTG 285 Leu Glu Val Pro Trp Leu Ser Ser Thr Val Ser Cys Ala Gln Gly Leu -15 -10 AGA TTG GCA CAA CAC AGA GTG CCT TTC TTT TAT TCA AAT GTC TCA TTA 333 Arg Leu Ala Gln His Arg Val Pro Phe Phe Tyr Ser Asn Val Ser Leu TGC AAA TTA TTG CTG CCA GCC AMM CTG CAC GGG 366 Cys Lys Leu Leu Pro Ala Xaa Leu His Gly 20 25

#### (2) INFORMATION FOR SEQ ID NO: 86:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 437 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 123..209
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5.7

seq SPAFLAVAGPGWA/RP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 86:

										0,						
GCTTTAATTG ATCCTGCTGC CCTCTGGGAG CACCCACACA AGCCAGGCTC TCACTGGACC 120																
TC														AGC . Ser		167
TGC Trp	AGT Ser	CCT Pro	GCC Ala	TTC Phe	: Lei	r GCA ı Ala	A GTO	G GCC L Ala	GGC Gly	/ Pro	GGC GGC	TGG Trp	GCA Ala	CGT Arg	CCT Pro	215
GGC Gly	TGI Cys	C DCC	Leu	AGG Arg	ACC J Thi	AAG Lys	TAT Tyr	Asp	TC: Sei	CAC Glr	CTC Leu	GCC Ala 15	Arg	CAC His	CTC Leu	263
TTG	CAG Gln 20	Pro	CAC Glr	TTC Phe	CCT Pro	GGT Gly 25	/ Let	ACC Thr	CTT Let	GGG Gly	ACC Thr	Leu	GTG Val	CAA Gln	CCT Pro	311
GCC Ala 35	His	TGG Trp	GGC Gly	ATO Met	GG/ Gl <sub>2</sub>	/ Gly	GGC Gly	C ACA	GG/	4 GGG 7 Gly 45	/ Val	TTC Leu	GGC Gly	GAG Glu	GGA Gly 50	359
					Ala					Cys				TGC Cys 65		407
				Xaa		GTC Val			ı Ala							437
(2)	(2) INFORMATION FOR SEQ ID NO: 87:  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 437 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain  (ix) FEATURE: (A) NAME/KEY: sig_peptide (B) LOCATION: 63116 (C) IDENTIFICATION METHOD: Von Heijne matrix															
	(	(xi)	SEQU	JENCE	E DES	SCRIE	PTIO	1: SE		=			GC/H	G		
AAC	CTGTC	GTC	ACA	rccci	rca i	AAAG1	rgaa(	CA G	rcgc	CATC	G GAG	GGCG1	TTTG	GAGG	AGACCG	60
TG														CCC Pro -5		107

GGG Gly								155
CCA Pro 15								203
GAC Asp								251
 GTT Val	 	299						
AAG Lys								347
CAA Gln								395
 AAG Lys 95	 	 	 	 	 	 		437

## (2) INFORMATION FOR SEQ ID NO: 88:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 113 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 3..62
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5.6

seq SLVCLLAMGKGLG/SS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 88:

AC ATG TAT TCC CAT CCC GTG TCC TCA CTG GTG TGT CTC CTG GCC ATG

Met Tyr Ser His Pro Val Ser Ser Leu Val Cys Leu Leu Ala Met

-20 -15 -10

GGC AAG GGA CTC GGG TCA TCC CAG GCC CTG GTC CAG CCA GAC ACC TGG
Gly Lys Gly Leu Gly Ser Ser Gln Ala Leu Val Gln Pro Asp Thr Trp

V	VO 99/0655	2		•	69	•	1	PCT/IB98/01236
-5			1		5		10	
	CAC ACC His Thr							113
(2)	INFORMAT	ION FOR	SEQ ID N	0: 89:				
	(i) SE	(A) LENG (B) TYPE (C) STRA	CHARACTER TH: 362 C: NUCLEI ANDEDNESS DLOGY: LI	base pai C ACID : DOUBLE	rs			
	(ii) M	OLECULE	TYPE: CD	NA				
	(vi) C		SOURCE: ANISM: Hos SUE TYPE:	-	ns			
		(B) LOCA (C) IDEN (D) OTHE	R INFORM	191 ON METHO ATION:	D: Von H score 5. seq FIFM	EVLGSGAFS		
			DESCRIPT	•				
							AG TCCCTGG	
CCTC	AGAAGC 1	TCAACTC	rg gaggca				GAT GAC TO Asp Asp Cy	
Ser		Lys Lys		Thr Asn		Lys Thr B	TTC ATT TT Phe Ile Pho	
							CTG GTG AAG Leu Val Ly: 5	
							AAG AAG TC Lys Lys Se 20	
							STG TTG AA Val Leu Ly	
							TAT GAG AG Tyr Glu Se	
	CAA GGG Gln Gly							362

(2) INFORMATION FOR SEQ ID NO: 90:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 384 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: CDNA	
<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F) TISSUE TYPE: Brain</pre>	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 241327     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 5.6</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 90:	
AAGCGGCGCA CCGGGHGAAG ATGGCGTTGG AGGTCGGCGA TATGGAAGAT GGGCAGCTTT	60
CCGACTCGGA TTCCGACATG ACGGTCGCAC CCAGCGACAG GCCGCTGCAA TTGCCAAAAG	120
TGCTAGGTGG CGACAGTGCT ATGAGGGCCT TCCAGAACAC GGCAACTGCA TGTGCACCAG	180
TATCACATTA TCGAGCTGTT GAAAGTGTGG ATTCAAGTGA AGAAAGTTTT TCTGATTCAG	240
ATG ATG ATA GCT GTC TTT GGA AAC GCA AAC GAC AGA AAT GTT TTA ACC Met Met Ile Ala Val Phe Gly Asn Ala Asn Asp Arg Asn Val Leu Thr -25 -20 -15	288
CTC CTC CCA AAC CAG AGC CTT TTC AGT TTG GCC AGA GCA GTC AGA AAC Leu Leu Pro Asn Gln Ser Leu Phe Ser Leu Ala Arg Ala Val Arg Asn -10 -5 1	336
CAC CTG TTG CTG GAG GAA AGA AGA TTA ACA ACA TAT GGG GTG CTG TGC His Leu Leu Glu Glu Arg Arg Leu Thr Thr Tyr Gly Val Leu Cys 5 10 15	384
(2) INFORMATION FOR SEQ ID NO: 91:  (i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 314 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR	

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

71

(ix)	FEATURE:	
LIX	, realure:	,

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 141..197

(F) TISSUE TYPE: Brain

- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 5.6

seq LVVTAWFFGMCRS/KA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 91:

ATCCCAGAAC ACCATTGGGA GAACGCCAGG ACACCGTGAA GGCTGAGCCG CCACTCGGTT 60

CTGATGCCGC ATCCATTGGT CAGTGCACGT TCTTTGAGCT TCCACTTGAG TGCACGTTCT 120

TTGAGCTTCC ACTTGAGTGC ATG TTC TTT GAG CTT CCA CTT GTA GTG ACT GCC Met Phe Phe Glu Leu Pro Leu Val Val Thr Ala -15

TGG TTC TTC GGG ATG TGC AGG AGC AAA GCG CTC TTA GGC AAT GCT CGT 221 Trp Phe Phe Gly Met Cys Arg Ser Lys Ala Leu Leu Gly Asn Ala Arg

TCT GCC CTG TGT TTA CAA ACC AAG GCC TGT GCC AGC TCT ACT CAG CCT 269 Ser Ala Leu Cys Leu Gln Thr Lys Ala Cys Ala Ser Ser Thr Gln Pro

GAC ACC CAT AAT GAG CAC CAT CCC AGG AAT CCC TGT CCC TAC TTG 314 Asp Thr His Asn Glu His His Pro Arg Asn Pro Cys Pro Tyr Leu 25

## (2) INFORMATION FOR SEQ ID NO: 92:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 316 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 155..286
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5.5

seg FLLIVANVHFSOT/WV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 92:

ATAAATAGCA TTGTTTACAT CGACCAAATA TTGCCTGTTT CCTTTAATTC AAATGCATTA GGTTCCCGCC TCCCTCTCT TCCCCGCCAT GTTGCTGTTT TAAGGCTTCA TATGTATTAA 120

CATTTCTCTG ATCAAAATTG TGGCTGTTTT CCTT ATG AAC CAT AAT ATA ATC ATT  Met Asn His Asn Ile Ile Ile -40	175
TGT GTG ATG TAC ATT GTG CCA TTT TTG ATG ACT AAA TGT CTA TAT TTC Cys Val Met Tyr Ile Val Pro Phe Leu Met Thr Lys Cys Leu Tyr Phe -35 -30 -25	223
TGC CAT TCC TGT AAG AGA GGG AGT TTT TTA CTG ATA GTA GCA AAT GTT Cys His Ser Cys Lys Arg Gly Ser Phe Leu Leu Ile Val Ala Asn Val -20 -15 -10	271
CAC TTC AGT CAA ACT TGG GTG TTC AGT GGT AAA CCA TAT AAA GGG His Phe Ser Gln Thr Trp Val Phe Ser Gly Lys Pro Tyr Lys Gly -5 1 5 10	316
(2) INFORMATION FOR SEQ ID NO: 93:  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 405 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain  (ix) FEATURE: (A) NAME/KEY: sig_peptide (B) LOCATION: 247309 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 5.5 seq LTGLCXCCLQALG/LA  (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 93:	
ATTGTCTTAG TCCCATCCTC TGTTCCCCTG GCCAGTTTGT CTAGCTGTGT GGTCTCTGTT	60
CTCTCCCTAC CGTGCCTTCC ATCCCAGCCA TCCCTGACTA CGTGTTTCCC CCACAGACAT	120
CACACTGGTT CACCTCGTTG ACCACCGTTT CCTTCTCCCC AAGTCTCCCG GGCAAGGGCT	180
GATTCTCCAG TCTCCTCTGG GAAGCTGGCC CTGAACCACT TAGAACCTAT CGCTCCTTCG	240
TCACCT ATG TCA TGT GGC AGC GCT GCC TCA CTT ACG GGT CTG TGT KSG Met Ser Cys Gly Ser Ala Ala Ser Leu Thr Gly Leu Cys Xaa -20 -15 -10	288
TGC TGC CTC CAA GCC CTG GGG CTT GCG TGG CGC CGT CGC GGT TTG ACG Cys Cys Leu Gln Ala Leu Gly Leu Ala Trp Arg Arg Gly Leu Thr -5 1 5	336
GGA CCG GGC CTC CCC CCT GTG TTG CAG ATA TGC TGT CCA AGG AGC CTC Gly Pro Gly Leu Pro Pro Val Leu Gln Ile Cys Cys Pro Arg Ser Leu	384

W	O 99/0655	52				73							PCT/IB98/01236			
10 15 20 CGT GGT GTG ACG GCT CCT ACT													25			
	GGT <b>GTG</b> Gly Val														405	
(2) 1	INFORMA	TION	FOR	SEQ	ID N	10: 9	94:									
	(i) S	(A) (B) (C)	LENG TYPE	TH: : NU ANDEC	302 ICLEI NESS	base C AC	pai CID OUBLE									
	(ii) !	MOLEC	CULE	TYPE	: CI	ANG										
	(vi) (	(A)		NISM	1: Ho		Sapie sin	ens								
	(ix)	(A) (B) (C) (D)	NAME LOCA IDEN OTHE	TION TIFI R IN	: 99 CATI	ON M	METHO ON:	D: V scor seq	e 5. VLFF	4 VGLI		trix A/MR				
AAAA	ATACCA (	GATGO	CACT	rc To	CAG	CTG	CAAT	raaci	ACT	ACT	TACTO	GA 1	'ACA'	TCAAA	60	
СССТО	CCAGAA '	CAAC	CAGT	TA TO	CAGGI	CAAC	C AAC	CAAGA			ln Al			AC AAC sp Asn	116	
CTC A Leu 1 -40	ACC TCT Thr Ser	Ala	Pro	Gly	Asn	Thr	Ser	Leu	Cys	Thr	Arg	GAC Asp	Tyr	Lys	164	
ATC A	ACC CAG Thr Gln	GTC Val	CTC Leu -20	TTC Phe	CCA Pro	CTG Leu	CTC Leu	TAC Tyr -15	ACT Thr	GTC Val	CTG Leu	TTT Phe	TTT Phe -10	GTT Val	212	
GGA (	CTT ATC Leu Ile	ACA Thr -5	AAT Asn	GGC Gly	CTG Leu	GCG Ala	ATG Met 1	AGG Arg	ATT Ile	TTC Phe	TTT Phe 5	CAA Gln	ATC Ile	CGG Arg	260	
	AAA TCA Lys Ser 10														302	
(2)	INFORMA	TION	FOR	SEQ	ID I	NO:	95:									

(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 99 base pairs
(B) TYPE: NUCLEIC ACID

WO 99/065	52	74	PCT	/IB98/
	(C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR	 E		
(ii)	MOLECULE TYPE: CDNA			
(vi)	ORIGINAL SOURCE: (A) ORGANISM: Homo Sapio (F) TISSUE TYPE: Brain	ens		
(ix)	FEATURE:  (A) NAME/KEY: sig_peptic (B) LOCATION: 1675  (C) IDENTIFICATION METHO (D) OTHER INFORMATION:	DD: Von Heijne ma		
(xi)	SEQUENCE DESCRIPTION: SEC	Q ID NO: 95:		
AGAAGTAGCC	GCAGG ATG GCG GCT AT Met Ala Ala Ala Me -20	G CSS TTG CTC TGG t Xaa Leu Leu Cy: -15	C TCG TCC TGT s Ser Ser Cys -10	51
GC TCC TGG Cys Ser Trp	GGC CCG GCG GCT GGT GCC Gly Pro Ala Ala Gly Ala -5	TTG CAG AAC CCC Leu Gln Asn Pro 5	CAA CGC GGG Gln Arg Gly	99
(2) INFORMA	TION FOR SEQ ID NO: 96:			
(i) S	EQUENCE CHARACTERISTICS:  (A) LENGTH: 485 base particles: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR			
(ii)	MOLECULE TYPE: CDNA			
(vi)	ORIGÎNAL SOURCE: (A) ORGANISM: Homo Sapi (F) TISSUE TYPE: Brain	ens		
(ix)	FEATURE:  (A) NAME/KEY: sig_peptice (B) LOCATION: 396470 (C) IDENTIFICATION METHOLOGY (D) OTHER INFORMATION:	DD: Von Heijne ma		
(xi)	SEQUENCE DESCRIPTION: SE	Q ID NO: 96:		
ATTTCTGCCC	ACGGGCATAA GTTCAAAAGA AA	GCTGCGAA AAGTTGG	AGA CTGCTGATGA	60
AACCAGTCAT	CTCCAGCCAC TCAACAAGCG TC	AGAGGACA AGCTCTG	TGG TGGAAGAGCA	120
TTTCCAAGCC	TCAGTATCTC CCACTGAAGC CG	CACCCCCT GCCACAG	GAG ACCAGAGTCC	180

TGGCCTGGGC ACCCAGCCAA AGCTGCCATC CAGCAGTGGC CTTCCTGCTG CAGACGTGTC 240

CCC.	rgcc <i>i</i>	ACA (	GCTG!	AAGAC	sc co	TTG	CACC	TTO	CCAC	ACCC	ACC	CGCCC	GC (	CTCC	CTTCAC	300
CCG	AGGGG	GA (	CTCC	GCTC	C TO	CTCCT	TTC	ATO	CCAT	GGAG	GAGO	CCA	GAC 1	rggto	GCCAC	360
AGT	GAAA	GAS (	CAAAT	racco	T GT	GCT	SAAGG	AC(	i		GAC 1 Asp 1			Lys 1		413
										Val	CTT Leu					461
	CAG Gln															485
(2)	INFO	RMAT	TION	FOR	SEQ	ID N	10: 9	7:								
	(i	.) SE	(A) (B) (C)	LENG TYPE STRA	TH: : NU NDED	283 CLEI NESS	DASE C AC C DO NEAR	pai ID UBLE								
	(i	i) M	OLEC	ULE	TYPE	: CD	NA									
	( v	·i) C	(A)		NISM	: Ho	mo S Bra	-	ens							
	(i	ж) <b>Е</b>	(B) (C)	NAME LOCA IDEN	TION	: 59 CATI	g_pe 14 ON M	2 ETHC	D: N	re 5	Heijn .3 CAFSL					
	( x	:i) S	SEQUE	NCE	DESC	RIPT	:NOI	SEC	Q ID	NO:	97:					
AGT	STGTO	SAA C	GCGT#	ACCTA	AR GO	CGGC	SAGGO	GAC	CATG	GWGA	CAGO	GGC	GY (	CGWG	CTGT	58
											TGG Trp					106
											GCA Ala					154
											GCT Ala					202
										Lys	CCT Pro					250
ccc	GTC	AGG	ATG	ATG	AGA	AGA	GAA	GGC	AGY	KGA						283

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Pro Val Arg Met Met Arg Arg Glu Gly Ser Xaa 40 45

(	2)	INFORMATION	FOR	SEO	TD	NO:	98 .

ı	•	١.	SECTIENCE	CHARACTERISTICS:	
1	. 1	,	SECUENCE	CHARACIERISTICS	

- (A) LENGTH: 390 base pairs
- (B) TYPE: NUCLEIC ACID
- (C) STRANDEDNESS: DOUBLE
- (D) TOPOLOGY: LINEAR

# (ii) MOLECULE TYPE: CDNA

- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain

### (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 286..333
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 5.3

seq CAVSLTTAAVAFG/DE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 98:

ACTNTGTGCT GGTGCTGCCA AAGTTTGTGA TTTTAAGAAA TTCTGCTGTG CTCTCCAGCA 60

CTGCGAGCTT CTGCCTTCCC TGTAGTTTCC CAGATGTGAT CCAGGTAGCC GAGATTCCGC 120

TGCCCGTGCT TCGGTAGCTT AAGTCTTTGC CTCAGCTTTT TTCCTTGCAG CCGCTGAGGA 180

GGCGATAAAA TTGGCGTCAC AGTCTCAAGC AGCGATTGAA GGCGTCTTTT CAACTACTCG 240

ATTAAGGTTG GGTATCGTCG TGGGACTTGG AAATTTGTTG TTTCC ATG AAA TCC TGC 297

Met Lys Ser Cys -15

GCA GTG TCG CTC ACT ACC GCC GCT GTT GCC TTC GGT GAT GAG GCA AAG A1a Val Ser Leu Thr Thr Ala Ala Val Ala Phe Gly Asp Glu Ala Lys

AAA ATG GCG GAA GGA AAA GCG AGC CGC GAG AGT GAA GAG GAG ACG
Lys Met Ala Glu Gly Lys Ala Ser Arg Glu Ser Glu Glu Glu Thr

5 10 15

### (2) INFORMATION FOR SEQ ID NO: 99:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 254 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA

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<ul><li>(vi) ORIGINAL SOURCE:</li><li>(A) ORGANISM: Homo Sapiens</li><li>(F) TISSUE TYPE: Brain</li></ul>
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 138200     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 5.2</pre>
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 99:
ATGTGATTWT KTTCCTATTT ATTTTAAATA CACACCCCA CAGGGCTCTG CCCCTGTAAA 60
AGAAAAAAA TCAAAACAAA CAAATAAATA ACCCCAAAGA GATGGACCCA GGGGAGAACG 120
CGTAAGTRTG AAGGGGC ATG AGT ATA CAC GAG TGT GCG TGT CTT TCC CTC  Met Ser Ile His Glu Cys Ala Cys Leu Ser Leu  -20 -15
TCC CTT ATT TGT CTC CGT ATG AGT CTC TCC TTG TAC CCT CCC CCT GCC  Ser Leu Ile Cys Leu Arg Met Ser Leu Ser Leu Tyr Pro Pro Ala -10 -5 1 5
TCG ATG ATA TTA CTC CCC CAG ACT TGG AAG CCG CGC Ser Met Ile Leu Leu Pro Gln Thr Trp Lys Pro Arg 10 15
(2) INFORMATION FOR SEQ ID NO: 100:  (i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 303 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR
(ii) MOLECULE TYPE: CDNA
<ul><li>(vi) ORIGINAL SOURCE:</li><li>(A) ORGANISM: Homo Sapiens</li><li>(F) TISSUE TYPE: Brain</li></ul>
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 178222     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 5.2     seq SGLSFLSVFSLWC/EP</pre>
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 100:
AAGATCTRGA ARCAGTRACC CTCTCTCTTT GCATRAGTTT CTCTTTTTTC TCTGAGTTAC 60
AGTTTTGARR RCAGCWRCTA ATTTTTTTAA TCCCTCGAAT AACTCAGTTT TAGGAACATT 120
CGCTCTCCCT AAGCCTTACC TTGAAACCAG TGTAGGATTT TGCTGCCACC CCGGAAG 17

					AGC Ser -10											225
					CTG Leu											273
					GCC Ala											303
(2)	INFO	RMAT	NOI	FOR	SEQ	ID N	10: 1	01:								
	(i	.) SE	(A) (B) (C)	LENG TYPE STRA	CHARA TH: : NU NDED LOGY	380 CLEI NESS	base C AC : DO	pai ID UBLE								
	(i	.i) M	OLEC	ULE	TYPE	: CE	NA									
	(1	i) C	(A)	ORGA	SOUR NISM UE T	: Ho			ns							
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 120374     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 5.2</pre>																
	(>	i) S	EQUE	NCE	DESC	RIPT	'ION:	SEC	) ID	NO:	101:					
AACI	TTG#	ATG (	SAATO	CAAAC	G TO	CATGO	GCGC	CAC	GGC7	AGCT	GTTC	CCAC	CAT C	CAGO	TGGGG	60
GCC	CGCC	CTT 7	rgtto	CACAC	CC TA	CAT	CAAC	G GAJ	ATTC	rgtt	GGGG	CTCAT	CAA T	TCGT	TGTG	119
ATG Met -85	GGG Gly	TTA Leu	AAG Lys	GAC Asp	AAA Lys -80	TCT Ser	CAG Gln	GCC Ala	CCC Pro	GCC Ala -75	TCA Ser	GGA Gly	CTG Leu	GGA Gly	GTT Val -70	167
					TCG Ser											215
					GAA Glu											263
					AGA Arg											311
		CAA Gln			CGG Arg											359

-20			-15	-10	
	 	 CGA Arg 1		31	80

### (2) INFORMATION FOR SEQ ID NO: 102:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 265 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 65..193
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5.2

seq PMQLLQVLSDVLA/EI

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 102:

# AATGCCAGTA TTAAGGATTT TTTTTCTAT TTTTACTCTT TAGTTAAAAT TATAAGACCT 60 AATT ATG AGT GAT CAA ATT AAA TTC ATT ATG GAC AGT CTC AAT AAG GAG Met Ser Asp Gln Ile Lys Phe Ile Met Asp Ser Leu Asn Lys Glu -40 -35 -30 CCC TTT AGG AAG AAC TAT AAT TTA ATC ACG TTT GAT TCC TTG GAG CCA Pro Phe Arg Lys Asn Tyr Asn Leu Ile Thr Phe Asp Ser Leu Glu Pro -25 -20 -15 ATG CAA CTA TTA CAA GTT CTC AGT GAT GTT CTG GCT GAG ATT GAC CCA Met Gln Leu Leu Gln Val Leu Ser Asp Val Leu Ala Glu Ile Asp Pro -10 -5 1

AAG GTA AGA GTT TTC TCT TTC TTT TTG ATG GGT AGC AGA AAG CCA ATT

Lys Val Arg Val Phe Ser Phe Phe Leu Met Gly Ser Arg Lys Pro Ile

5 10 15 20

TCT CCC TCT TGG 265
Ser Pro Ser Trp

(2) INFORMATION FOR SEQ ID NO: 103:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 354 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE

(D) .TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide
(B) LOCATION: 25..336

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 5.1

seq SSVASLTATPSLA/SP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 103:

AAACCTTGTA C		ATG TCT CCT TCC Met Ser Pro Ser		
CTG TGG TCA . Leu Trp Ser ! -95	ATG TGT CTG GAG Met Cys Leu Glu -90	GTC CCC TCC TTT Val Pro Ser Phe	Thr Ala Thr As	C TCA 99 p Ser -80
GTG AAC TGC (Val Asn Cys	GGC TGC TGT TTG Gly Cys Cys Leu -75	GAG CTC GCG ACG Glu Leu Ala Thr -70	GAG CCG GCT CG Glu Pro Ala Ar	g Asn
Ile Arg Ser	ACC ACC AGG GCT Thr Thr Arg Ala -60	TCT CTG CTG AGG Ser Leu Leu Arg -55	G TGC AGC TCA TT g Cys Ser Ser Ph -50	C ACT 195 e Thr
TCA ACC AGG : Ser Thr Arg : -45	AAC TCT ACG GGA Asn Ser Thr Gly	ATT TCA GCG CTG Tle Ser Ala Leu -40	G CCT CCC GCG GC Pro Pro Ala Al -35	C CCA 243 a Pro
ATG GCC TGG Met Ala Trp -30	CCA TTC TCA GCC Pro Phe Ser Ala -25	TCT TTG TCA ACG	G TTG CCA GTA CC Leu Pro Val Pr -20	T CTA 291 o Leu
ACC CAT TCC Thr His Ser	TCA GTC GCC TCC Ser Val Ala Ser -10	TTA ACC GCG ACA Leu Thr Ala Thr -5	Pro Ser Leu Al	A TCT 339 a Ser 1
CCT ACA AGA . Pro Thr Arg				354

# (2) INFORMATION FOR SEQ ID NO: 104:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 226 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA

<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F) TISSUE TYPE: Brain</pre>	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 155202     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 5.1</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 104:	
AACTCTGGAA TGAAGGGGGG AATTACGGGT TGGTGGTCGG CTTCATGTTA GGAGGACACC	60
CCTCCATCTG TTCACAGCTC AGCCTGTTTC CAATTTAAAG CCCAGAAGAA GCCTTCCCAG	120
CCTACTCAGA ATCCCACATC CTCTCCTCTC TCTT ATG GAT CTC AGT TTT CAT TTA  Met Asp Leu Ser Phe His Leu -15 -10	175
TTA CTA GAT CCT TCC TCT ACT CAA TCA AGC ATA CTG AAG CAC CTC CCA Leu Leu Asp Pro Ser Ser Thr Gln Ser Ser Ile Leu Lys His Leu Pro -5 1 5	223
TGT Cys	226
(2) INFORMATION FOR SEQ ID NO: 105:  (i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 447 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE:  (A) ORGANISM: Homo Sapiens  (F) TISSUE TYPE: Brain  (ix) FEATURE:  (A) NAME/KEY: sig_peptide  (B) LOCATION: 289366	
(C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 5 seq VISVLILVGFGAC/IY  (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 105:	
(XI) SEQUENCE DESCRIPTION: SEQ ID NO: 105:	
ATCTCTGATG GGCAGGGAGA GATACCAGGG TGCTGAGCCA GTCCAGGACT GCCCCCTCCT	60
GGCCCACTCA GAGCCCCTGG GTGTGAGAAG CTCGTCTCCC GTGGGTTGCA TTGGCTCTGC	120
CCTATCTCTG CCTCCAGCAC CCAGGGCGGC CGCAGATGGC AGTGTCTCTG GGGACAGCAG	130

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/00332			

CTGCGAATGA GTCCACGGGC CAACGCTGAG CTGCTCAGGC TGAGGCGGTG TGCTCAGCAC	240									
AGAGCCCCCG GAACTGGCAT CTGCAGGGCG TGAGCCAARG CCGCCGCG ATG CCG CAC  Met Pro His  -25	297									
TTC CTG GAC TGG TTC GTG MCG GTC TAC TTG GTC ATC TCG GTC CTC ATT Phe Leu Asp Trp Phe Val Xaa Val Tyr Leu Val Ile Ser Val Leu Ile -20 -15 -10	345									
CTG GTG GGC TTC GGC GCC TGC ATC TAC TAC TTC GAG CCG GGC CTG CAG Leu Val Gly Phe Gly Ala Cys Ile Tyr Tyr Phe Glu Pro Gly Leu Gln -5 1 5	393									
GAG GCG CAC AAG TGG CGC ATG YAG CGC CCC TGG TGG ACC GCG ACC TCC Glu Ala His Lys Trp Arg Met Xaa Arg Pro Trp Trp Thr Ala Thr Ser 10 20 25	441									
ACT GGG Thr Gly	447									
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 195 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE:  (A) ORGANISM: Homo Sapiens  (F) TISSUE TYPE: Brain  (ix) FEATURE:  (A) NAME/KEY: sig_peptide  (B) LOCATION: 79168  (C) IDENTIFICATION METHOD: Von Heijne matrix  (D) OTHER INFORMATION: score 5  seq IVGLLAQLEKINA/EP										
AACAGAAAGA TGACTTAAAG AACGTGGGGA GTCGCTCGCA GTTCGATTAT CTGCAATTAT	60									
GAAATGAAGT AACTCAAG ATG AGC AAG TTA AAA GTG ATA CCA GAA AAA AGC  Met Ser Lys Leu Lys Val Ile Pro Glu Lys Ser  -30 -25 -20	111									
CTT ACC AAT AAT TCT AGG ATC GTA GGA CTC CTG GCT CAA CTG GAG AAG Leu Thr Asn Asn Ser Arg Ile Val Gly Leu Leu Ala Gln Leu Glu Lys -15 -10 -5	159									
ATC AAT GCT GAG CCT TCA GAA TCW GAC ACT AGC CGG Ile Asn Ala Glu Pro Ser Glu Ser Asp Thr Ser Arg  1 5	195									

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(ii) MOLECULE TYPE: CDNA

(A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain

(A) NAME/KEY: sig\_peptide

(vi) ORIGINAL SOURCE:

(ix) FEATURE:

(2)	INFO	RMAT	ION	FOR	SEQ	ID 8	10: 1	107:							
	(i)	) SE	(A) (B) (C)	ICE C LENG TYPE STRA TOPO	TH: : NU NDEC	166 CLEI NESS	base C AC : DC	pai ID UBLE							
	(i:	i) M	OLEC	ULE	TYPE	: C	NA								
	(v:	i) O	(A)	NAL ORGA TISS	NISM	1: Hc			ens						
	(i:	х) F	(B) (C)	IRE: NAME LOCA IDEN OTHE	TION	1: 38 CATI	010 ON M	6 1ETHC	D: V	re 5	leijn MAFI				
	(x:	i) S	EQUE	NCE	DESC	CRIPT	ION:	SE(	) ID	: ОИ	107:	;			
AAC'	rgctg(	CT C	CACAC	GAAGO	CA G	rgago	GATG	A TGO	CCAGO				a Sei	G CGC r Arg	
	GCT (														103
	GAA . Glu 1														151
	TGC Cys														166
(2)	INFO			FOR											
			(A) (B) (C)	TYPE STRA	STH: E: NO ANDE	278 JCLE DNES	base IC AG S: DG	e pa: CID DUBL							

(B) LOCATION: 84..230

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.9

seq VTVCCXLVAFLFC/IL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 108:

AARAGACTCC GCCCCCTTCC TTGGAGCGCC GCGNCTCGGG CTGAGGGAGC TCGGGCCAAT 60

CAGAGGGACG GCCCCAGART GGC ATG GTA GAT GGA ACG CAG CTG AGA GGT CTG 113

Met Val Asp Gly Thr Gln Leu Arg Gly Leu

-45

ACA AGA ATG TAC CAG GTC CCA CTA MCA CTG GAT CGG GAT GAG ACC CTG

Thr Arg Met Tyr Gln Val Pro Leu Xaa Leu Asp Arg Asp Glu Thr Leu

-35

-30

-25

GTA CGG CTC CGC TTC ACC ATG GTG GCC CTG GTC ACG GTC TGC TGT MCA 209
Val Arg Leu Arg Phe Thr Met Val Ala Leu Val Thr Val Cys Cys Xaa
-20 -15 -10

CTT GTC GCC TTC CTC TGC ATC CTC TGG TCC CTG CTC TTC CAC TTC

Leu Val Ala Phe Leu Phe Cys Ile Leu Trp Ser Leu Leu Phe His Phe

-5

1

257

AAG GAG ACA ACG GCC ACA GGG Lys Glu Thr Thr Ala Thr Gly 10 15

278

# (2) INFORMATION FOR SEQ ID NO: 109:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 217 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 116..193
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.9

seg LISMLQMLAVIIT/NT

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 109:

ACTTGAAGTT CYYCTGGGGG CAATGAGATG GCAGCTATAC AGCGAGTCTG AAAAGAACAT 60

CCACATTCCT AATCCCTAGG AATATGATTA TTGGAAAATA GATATAATTA TACAA ATG 118 Met

-25	Gln	AAC Asn	TTC Phe	CTT Leu	GTT Val -20	CTC Leu	AAC Asn	AGT Ser	GTC Val	TGG Trp -15	TAC Tyr	CTA Leu	ATA Ile	AGC Ser	ATG Met -10	166
		ATG Met														214
GGG Gly																217
(2)	INFO	ORMAT	NOI	FOR	SEQ	ID N	10: 1	10:								
	į)	.) SE	(A) (B) (C)	ICE C LENG TYPE STRA TOPO	TH: : NU NDED	426 CLEI NESS	base C AC : DO	pai ID UBLE								
	( i	.i) M	OLEC	ULE	TYPE	: CD	AA									
	(1	ri) C	(A)	NAL ORGA TISS	NISM	: Ho			ns							
	<b>(</b> i	.×) F	(A) (B)	RE: NAME LOCA IDEN	TION	: 55	23	1					<b></b>			
				OTHE					scor	e 4.	_					
	(;)	(i) S	(D)	OTHE	R IN	FORM	ATIC	)N:	scor seq	e 4. LVEM	9 ICLEV	LGSS				
AAT:		(i) S	(D) SEQUE	OTHE	R IN	FORM	ATIC	SEC	scor seq	e 4. LVEM	9 ICLEV	LGSS	A/GE	)	ATG Met	57
GAA	CAG!		(D) SEQUE TAGAA	OTHE :NCE AAAC# AGT	R IN  DESC  AA AC  TCT	FORM RIPT CCAGI	TAGAT	SEC T TTT	scor seq DID	e 4. LVEM NO: GAAC	9 ICLEV 110: AAAA	LGSS	A/GE	AG.A.A	Met TCA	57 105
GAA Glu CTT	TGT Cys GTG	AAT T	(D) SEQUE TAGAL AAT Asn -55 GCT	OTHE NCE AAACA AGT Ser TCT	DESC AA AC TCT Ser	FORM RIPT CCAGT TTA Leu	TAGAT  AAA  Lys	SEC TTTT AAG Lys -50 GCT	scor seq ID TTTTC TGT Cys	e 4. LVEM NO: GAAC TTA Leu CAA	9 ICLEV 110: AAAA CTA Leu ACT	AATCT GTT Val	GAA Glu -45 GCA	AGAA AAG Lys	Met TCA Ser	-
GAA Glu CTT Leu AAG	TGT Cys GTG Val	CAA Gln AAA Lys	(D) SEQUE TAGAF AAT Asn -55 GCT Ala	OTHE CNCE AAACA AGT Ser TCT Ser	DESC AA AC TCT Ser TAT Tyr	TTA Leu GAA	AAA Lys ATT Ile -35 GAA	SECT TTTT  AAG Lys -50 GCT Ala TTA	scor seq ) ID TTTTC TGT Cys TTC Phe	e 4. LVEM NO: GAAC TTA Leu CAA Gln	9 ICLEV 110: AAAA CTA Leu ACT Thr	GTT Val GCT Ala -30	GAA Glu -45 GCA Ala	AGAA AAG Lys AGC Ser	Met TCA Ser AAG Lys	105
GAA Glu CTT Leu AAG AAG	TGT Cys GTG Val CCA Pro -25	CAA Gln AAA Lys -40	(D) SEQUE TAGAF AAT Asn -55 GCT Ala TCB Ser GAA	OTHE CNCE AAACA AGT Ser TCT Ser ATT Ile	DESC AA AC TCT Ser TAT Tyr GCT Ala	TTA Leu GAA Glu -20 GGT	TAGAT  AAA Lys  ATT Ile -35 GAA Glu  TCA	SECT TTTT  AAG Lys -50 GCT Ala TTA Leu AGT	scor seq PID TTTTC TGT Cys TTC Phe ATT Ile	e 4. LVEM NO: GAAC TTA Leu CAA Gln AAA Lys	9 ICLEV 110: AAAA CTA Leu ACT Thr CCA Pro -15 GAC	GTT Val GCT Ala -30 TAT Tyr	GAA Glu -45 GCA Ala TTA Leu	AGAA AAG Lys AGC Ser GTA Val	Met TCA Ser AAG Lys GAA Glu ACT	105

GCA GAC ATT GAA GAC CAG CTG ATT CAA AAG GTC AGA GAG TCA AAG TGG 345

	WO 3	77/003	32						8	6			•		•	011127-
Ala	Asp	Ile. 25	Glu	Asp	Gln	Leu	Ile 30	Gln	Lys	Val	Arg	Glu 35	Ser	Lys	Trp	
TTT Phe	GCC Ala 40	CTT Leu	CAG Gln	ATA Ile	GAT Asp	GAG Glu 45	TCA Ser	TCA Ser	GAA Glu	ATC Ile	TCA Ser 50	AAT Asn	ATC Ile	ACA Thr	CTT Leu	393
CTT Leu 55	TTG Leu	TGC Cys	TAT Tyr	ATT Ile	CGT Arg 60	TTC Phe	ATT Ile	GAT Asp	TAT Tyr	GAT Asp 65						426
(2)			EQUEN (A) (B)	CE C LENG TYPE	CHARA STH:	ACTER 95 b	NO: : RISTI Dase IC AC	ICS: pair								
							NEAF	₹								
				NAL ORG <i>A</i>	SOUF MSINA	RCE: i: Ho	omo S Bra	•	ens							
	<b>( )</b>	LX) I	(B) (C)	NAME LOCA IDEN	TION TIFI	: 15 CATI	ig_pe 583 ION M	3 1ETHO	D: V	e 4.	8		itrix 'C/KK			
	()	<i) 5<="" td=""><td>SEQUE</td><td>ENCE</td><td>DESC</td><td>CRIPT</td><td>CION:</td><td>: SE(</td><td>) ID</td><td>NO:</td><td>111:</td><td>:</td><td></td><td></td><td></td><td></td></i)>	SEQUE	ENCE	DESC	CRIPT	CION:	: SE(	) ID	NO:	111:	:				
ATTO	Jaaai	AAT i	AAAA				AGT Ser -20									50
TGG Trp	CTT Leu -10	GTT Val	GCT Ala	CTT Leu	TTG Leu	GAG Glu -5	ATG Met	TGT Cys	GTG Val	TGT Cys	AAG Lys 1	AAG Lys	TCC Ser	AGG Arg		95
(2)	INF	ORMA'	TION	FOR	SEQ	ID I	NO:	112:								
	(:	i) S!	(A) (B) (C)	LENC TYPE STRA	GTH: E: NU ANDEI	473 JCLE DNES	RIST: base IC AC S: DC	e pa: CID DUBLE								
	(:	ii)	MOLE	CULE	TYP	E: C	DNA									•
	(	vi) (		ORG	ANISI	4: H	omo : : Bra		ens							

WO 99/06552 87

ix)	FEATURE:
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(A) NAME/KEY: sig\_peptide (B) LOCATION: 342..395

(C) IDENTIFICATION METHOD: .Von Heijne matrix

(D) OTHER INFORMATION: score 4.8

seq LEAISSLSSFVLG/RM

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 112:

ACTGTTTATA GATATTTTGT TTCCCTGAGC AACAGAAAAT GCAGTGCTTA TTTAACTTAG CAGCAATGCC TTGTAAAAAT ATAAGCCTGC AGATGGCAAT GGCCTCTATT TTTCTTCCAC 120 AAGTTTCTTC CAATTCAGAG CCCGTGCCTT CCTTCAGCCA CAGAGCGCAC AACAGCATGG 180 ATGAGATTGA GTCAGCCCTC TTACATTGTT GGCCTACAGC TATGGAGCTA CCTTTGCAGA 240 GTTGTCCACT TTGGGGTTTG AGCATGGGAA GTAAATTCAG AGATGCAAGT ATCTGGGAGA 300 GGGCATGAAC TCGTGAGAAA GTCCTCATAT TCTGAGCTCC T ATG ACA GTT TTG CCT 356 Met Thr Val Leu Pro TTA GAA GCT ATC TCG TCT CTT AGC AGC TTT GTT TTG GGC AGA ATG AAT 404 Leu Glu Ala Ile Ser Ser Leu Ser Ser Phe Val Leu Gly Arg Met Asn AGC AGA GGG GCA GGA AAG ACC CAG AAT CTT GAT GCC AGC TCC YTG CTT 452 Ser Arg Gly Ala Gly Lys Thr Gln Asn Leu Asp Ala Ser Ser Leu Leu TTA CTC TGC TGC TTG ATA CTG 473 Leu Leu Cys Cys Leu Ile Leu 20

# (2) INFORMATION FOR SEQ ID NO: 113:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 386 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 12..101
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.8

seq ILFCVGAVGACTL/SV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 113:

AAT!	CAC	AGA A		: Gl			, Ser			, His	CTG Leu	50
		AAT Asn -15										98
		GTC Val										146
		ACC Thr										194
-		ACA Thr								 		242
		TTG Leu 50							-	 		290
		CTC Leu								 		338
		GAC Asp	-							 		386

# (2) INFORMATION FOR SEQ ID NO: 114:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 147 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 10..84
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.8

seq ALFYSVVVSTVSG/NE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 114:

AAGAGTCTG ATG AAT AGC AGT AAA GAA GAA ATG CGC GAA CTG GCA GCG TTG 51 Met Asn Ser Ser Lys Glu Glo Met Arg Glu Leu Ala Ala Leu

WQ 99/06552	00	PCT/IB98/01236

WO 99/06552						89							FC 1/1B30				
			-2	25			-20					- 2	-15				
			TCT Ser														99
	ATG Met	ATA Ile	GAA Glu	CAG Gln	CTT Leu 10	ATA Ile	AAG Lys	ACT Thr	ACA Thr	AAA Lys 15	GAC Asp	AAT Asn	CAC His	AGC Ser	CTA Leu 20	CGG Arg	147
	(2)	INFO	ORMAT	NOI	FOR	SEQ	ID N	NO: 1	L15:								
		(i	.) SE	(A) (B) (C)	LENG TYPE STRA	CHARA STH: C: NU ANDED OLOGY	297 ICLEI NESS	base C AC C DC	pai ID UBLE								
		(i	i) M	OLEC	CULE	TYPE	: CI	ONA									
		(7	7i) C	(A)	ORGA	SOUF NISM SUE T	: Ho		•	ens							
		(i	.x) E	(A) (B) (C)	NAME LOCA I DEN	C/KEY ATION STIFI CR IN	: 55 CATI	021 ON M	.0 1ETHC	D: V	e 4.	_					
		()	(i) S	EQUE	ENCE	DESC	RIPT	CION:	SEÇ	D ID	NO:	115	:				
	AAGT	TGTO	SCG (	CCGG1	rccc	rg go	CCT	GAGC	r cc	GCT	ccec	CTG	GGC	GCC 1	rgcg	ATG Met	57
			GAT Asp														105
			TCT Ser														153
			CGG Arg														201
	TCC	AGC	TGT	GCC	CCT	AGT	GTC	CAG	ATG	AAG	ATC	AAA	GAG	CTT	TAC	CGA	249

Ser Ser Cys Ala Pro Ser Val Gln Met Lys Ile Lys Glu Leu Tyr Arg

CGA CGC TTT CCC CGG AAG ACC CTG GGG CCC TCT GAT CTC TCC TCC GGG Arg Arg Phe Pro Arg Lys Thr Leu Gly Pro Ser Asp Leu Ser Ser Gly

20

15

297

(2) INFORMATION FOR SEQ ID NO: 116:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 141 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 1..87

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.7

seq LCYLSIFCLGVLF/II

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 116:

ATG CCT TGT ATA TCT CTC TTA GGT CTA CTT TAT AAT TTT GTT CAA GTC

Met Pro Cys Ile Ser Leu Leu Gly Leu Leu Tyr Asn Phe Val Gln Val

-25

CTC TGT TAC TTA TCG ATC TTC TGT CTA GGT GTT CTG TTC ATT ATT GAA

Leu Cys Tyr Leu Ser Ile Phe Cys Leu Gly Val Leu Phe Ile Ile Glu

-10

-5

CGT GGT TCA TTA AAA GTC TCC AAA TTA ATC TGT AGG CCA CCA GGG
Arg Gly Ser Leu Lys Val Ser Lys Leu Ile Cys Arg Pro Pro Gly
10
15

# (2) INFORMATION FOR SEQ ID NO: 117:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 307 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig peptide
  - (B) LOCATION: 167..211
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.7

seq IAVLFCFFLLIIF/QT

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 117:

AACT	'AAG'	WN !	KTTC	AGCA)	AA TA	ACTT	TCA	A CA	rtcc	CTTC	TGT	CCTT	rct :	TTGT'	AATTTT	60	
AGAA	AGCI	CT (	GATT	TGT	TT C	ATTT	CAGO	TG	GAGA	CTTA	AATO	SACA	CCA A	AGCA	AAGCCT	120	
ACTI	'AGT'I	TA (	GATCI	rccao	SA A	ATTGO	GCTG	TG(	GAAA!	AAA	TCA	t		AAG : Lys :		175	
			TTT Phe													223	
			GAA Glu													271	
			AGG Arg													307	

# (2) INFORMATION FOR SEQ ID NO: 118:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 396 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA-
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 253..381
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.7

seq STWSSASLRGSWQ/QG

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 118:

AATACTGATG	TCTYCCGAAA AACA	CAGCCC CAAGGGAGTC	GAGACGWTGT ACCAGGTAGA	60
ATAAGGCACA	GGGGAGCCGC TTGA	CAAATC AGACGACGGC	AGCCGGCCTG CCTGCCCGGT	120
ATGTGGCCAA	ATATGGGCGA GGCC	AAGGTT GGGGTGTGAA	AGTGCGTGAC GTTTACACCC	180
ACGTGGGCGT	CTGTGCACGT GCGT	GTGTGC GTGTGAGCTG	CCTGTGGGCA TCTGCAGAAG	240
CAGACATTCT			TT TTG GGT TCC AGG GTG al Leu Gly Ser Arg Val -35	291

ATG CCA GCG AGT TGT GTT TCT GAG AGA CGA AGG CCT TCC TTC CAG
Met Pro Ala Ser Cys Val Ser Glu Arg Arg Arg Lys Pro Ser Phe Gln

WO 99/06552					PCT/IB98/01236
-30	25	9	22		
	-25		-20	-15	
GTT TCC ACG TGG Val Ser Thr Trp					
ATG CCA GGC					396
Met Pro Gly 5					
(2) INFORMATION	FOR SEQ ID 1	NO: 119:			
(A) (B) (C)	CE CHARACTER LENGTH: 193 TYPE: NUCLEI STRANDEDNESS TOPOLOGY: LI	base pairs C ACID S: DOUBLE			
(ii) MOLEC	ULE TYPE: CO	ANC			
(A)	NAL SOURCE: ORGANISM: Ho TISSUE TYPE:				
(B) (C)	RE: NAME/KEY: si LOCATION: 14 IDENTIFICATI OTHER INFORM	13187 ION METHOD: MATION: Sco	Von Heijne π re 4.6 FLYLKSVFDVS		
(xi) SEQUE	NCE DESCRIPT	CION: SEQ ID	NO: 119:		
ATGGAGATCC ATAAC	ATGAT CCTAC	ATGAA TGTTTC	AATA TTGTAT	CCT GTAAGTTA	CT 60
TTTACATTGA CAGTT	CTGAA ATTCA	IGTTG AGTGTT	AATT AGGCAG	SAAA TCAGAAGG	GA 120
GGTTTTGTAG AAGGI		t Gly Phe Le		AA AGT GTT TI ys Ser Val Ph	
GAT GTA TCA TTG Asp Val Ser Leu -5					193
(2) INFORMATION	FOR SEQ ID	NO: 120:			
(A) (B) (C)	NCE CHARACTE LENGTH: 460 TYPE: NUCLE STRANDEDNES TOPOLOGY: L	base pairs IC ACID S: DOUBLE			
(ii) MOLEC	CULE TYPE: C	DNA			

(vi) ORIGINAL SOURCE:

(A)	ORGANISM:	Homo	Sapiens
(E)	TISSUE TY	PE: B	rain

# (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 254..436

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.6

seq LLLLHGGGHSALS/WA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 120:

AAGTTTACGG AGCCGGTGGG CGGTAGGCGG TGCTACGGGT AGCTGGGTGC TGTCCAAAGG	60
CGACAGGGCG TCGTTAGGGG AGCGAGTCGT GACCGGTTGG GCCACACTCA ACGTGGGACG	120
AAGCTTCGCC TACTGTTTGA CTACGTGCGT GCAGCCTCCC CTCGATGTCG GCCCTCGAAA	180
AGAGCATGCA CCTCGGCCGC CTTCCCTCTC GCCCACCTCT ACCCGGCAGC GGGGGCAGTC	240
AGAGCGGASC AAG ATG CGA ATG GGC CCT GGA AGA AAG CGG GAC TTT TCC  Met Arg Met Gly Pro Gly Arg Lys Arg Asp Phe Ser  -60 -55 -50	289
CCT GTT CCT TGG AGT CAG TAT TTT GAG TCC ATG GAA GAT GTA GAA GTA Pro Val Pro Trp Ser Gln Tyr Phe Glu Ser Met Glu Asp Val Glu Val -45 -40 -35	337
GAG AAT GAA ACT GGC AAG GAT ACT TTT CGA GTC TAC AAG AGT GGT TCA Glu Asn Glu Thr Gly Lys Asp Thr Phe Arg Val Tyr Lys Ser Gly Ser -30 -25	385
GAG GGT CCA GTC CTG CTC CTT CTG CAT GGA GGA GGT CAT TCT GCC CTT Glu Gly Pro Val Leu Leu Leu His Gly Gly Gly His Ser Ala Leu -15 -10 -5	433
TCT TGG GCT GTG TTC ACG GCA GCT ARG Ser Trp Ala Val Phe Thr Ala Ala Xaa	460

### (2) INFORMATION FOR SEQ ID NO: 121:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 275 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 207..245
  - (C) IDENTIFICATION METHOD: Von Heijne matrix

			<b>7</b> '
. (D)	OTHER	INFORMATION:	score 4.6 seq MIFLLYLLPSSEE/RR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 121:

AAACAGACAG GTATGGAGTC TGGGTGGGGC CACGTGTACC CTCCCATCCT TAGAAAGAGT 60

GTGACACCAA GGGACAGATG CTGGCGTASG CGGGTTTTGT TTTGGAGGGT TTTTTGTTTG 120

TTTTTACAAA AATTAAGATA TTTCTGAGTT TATTATGAGG CTTTTAGTTT TACAATCATA 180

CTAAAAAGATA ATTGTTCCTC TATAAA ATG ATT TTC CTT CTG TAC CTC TTG CCT 233

Met Ile Phe Leu Leu Tyr Leu Leu Pro -10 -5

TCT TCT GAA GAA AGG AGA AAA TTG CTT TTT AGT CCC CAC AGG 275

Ser Ser Glu Glu Arg Arg Lys Leu Leu Phe Ser Pro His Arg 10

# (2) INFORMATION FOR SEQ ID NO: 122:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 445 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 236..418
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.6

seq LLLLHGGGHSALS/WA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 122:

GCGGTAGGCG GGTGCTACGG GTAGCTGGGT GCTGTCCAAA GGCGACAGGG CGTCGTTAGG	60
GGAGCGAGTC GTGACCGGTT GGGCCACACT CAACGTGGGA CGAAGCTTCG CCTACTGTTT	120
GACTACGTGC GTGCAGCCTC CCCTCGATGT CGGCCCTCGA AAAGAGCATG CACCTCGGCC	180
GCCTTCCCTC TCGCCCACCT CTACCCGGCA GCGGGGGCAG TCAGAGCGGA SCAAG ATG Met	238
CGA ATG GGC CCT GGA AGA AAG CGG GAC TTT TCC CCT GTT CCT TGG AGT Arg Met Gly Pro Gly Arg Lys Arg Asp Phe Ser Pro Val Pro Trp Ser -60 -55 -50 -45	286
CAG TAT TTT GAG TCC ATG GAA GAT GTA GAA GTA GAG AAT GAA ACT GGC Gln Tyr Phe Glu Ser Met Glu Asp Val Glu Val Glu Asn Glu Thr Gly	334

									7.	,						
				-40					-35					-30		
AAG Lys	GAT Asp	ACT Thr	TTT Phe -25	CGA Arg	GTC Val	TAC Tyr	AAG Lys	AGT Ser -20	GGT Gly	TCA Ser	GAG Glu	GGT Gly	CCA Pro -15	GTC Val	CTG Leu	382
CTC Leu	CTT Leu	CTG Leu -10	CAT His	GGA Gly	GGA Gly	GGT Gly	CAT His -5	TCT Ser	GCC Ala	CTT Leu	TCT Ser	TGG Trp 1	GCT Ala	GTG Val	TTC Phe	430
		GCG Ala														445
(2)	(i) (i)	ORMAT	CQUEN (A) (B) (C) (D) MOLEC DRIGI	ICE ( LENG TYPE STRA TOPO	CHARACTH: CHARACTH: CHARACTHE CHARAC	ACTER 138 DCLEI DNESS : LI C: CE RCE:	RISTI base CC AC S: DC INEAF DNA	ICS: pai CID OUBLE	2							
		.×) E	(A) (B) (C) (D)	NAME LOCA I DEN OTHE	TION TIFI R IN	I: 49 CATI IFORM	O96 ON M	S METHO ON:	D: V scor seq	te 4.	5 JISII	ASIE	itrix PS/QE			
ATA															G AGC 1 Ser	57
CTA Leu	TTA Leu	AAT Asn	CTC Leu -10	ATT Ile	TCA Ser	ATC Ile	TTA Leu	GCA Ala -5	AGT Ser	ATT Ile	CCC Pro	AGT Ser	CAA Gln 1	TTT Phe	AAA Lys	105
		TTT Phe														138
(2)		DRMA:	EQUEI (A) (B)		CHARA STH: E: NU	ACTE: 94 k JCLE:	RIST: base EC AC	ICS: pain								

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

<pre>(vi) ORIGINAL SOURCE:    (A) ORGANISM: Homo Sapiens    (F) TISSUE TYPE: Brain</pre>	
(ix) FEATURE:  (A) NAME/KEY: sig_peptide  (B) LOCATION: 1185  (C) IDENTIFICATION METHOD: Von Heijne matrix  (D) OTHER INFORMATION: score 4.5  seq LMLLWPVHPLLVG/HR	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 124:	
AAGATTCATC ATG GGC ACC ACC TCC AAC ATG GTC ACC ACC ATC CAT CTC  Met Gly Thr Thr Ser Asn Met Val Thr Thr Ile His Leu  -25  -20 -15	)
ATG TTG CTG TGG CCA GTG CAT CCA TTA CTG GTG GGC CAC CGC GGG  Met Leu Trp Pro Val His Pro Leu Leu Val Gly His Arg Gly -10 -5 1	ī
(2) INFORMATION FOR SEQ ID NO: 125:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 481 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: CDNA	
<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F) TISSUE TYPE: Brain</pre>	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 41343     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 4.4     seq ISHILAFFAASDG/IV</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 125:	
AAGTTCCCTC AGCGCCCGTA GCTTCGGCGG AGTCTGCGCG ATG GGC GAC CCG GAA 59 Met Gly Asp Pro Glu -100	5
AGG CCG GAA GCG GCC GGG CTG GAT CAG GAT GAG AGA TCA TCT TCA GAC 103 Arg Pro Glu Ala Ala Gly Lou Asp Gln Asp Glu Arg Ser Ser Asp -95 -90 -85	3
ACC AAC CAA AGT GAA ATA AAG TCA AAT GAA GAG CCA CTC CTA AGA AAG 15: Thr Ash Glu Ser Glu Ile Lys Ser Ash Glu Glu Pro Leu Leu Arg Lys	1

					,	′				
-80			<del>-</del> 75			-70 ·			-65	
			GTC Val							199
			GCA Ala							247
			CTC Leu							295
			CAC His							343
			TTG Leu							391
			TTC Phe							439
		 _	AGT Ser				 			481

# (2) INFORMATION FOR SEQ ID NO: 126:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 197 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 3..50
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.4

seq GLFSLLPHPPCVG/RV

95

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 126:

AG ATG GAT GCA GGC TTA TTT TCT CTG CTT CCC CAT CCT CCA TGT GTT

Met Asp Ala Gly Leu Phe Ser Leu Leu Pro His Pro Pro Cys Val

-15

-10

-5

GGC AGG GTG CTG CCA CAG TCT AGG TAT CAT CTG CAT CCA AGA TCA CCT

'	VO 99	/0655	<b>i</b> 2						9	8					rc	1/10/0
Gly	Arg 1	Val	Leu	Pro	Gln 5	Ser	Arg	Tyr	His	Leu 10	His	Pro	Arg	Ser	Pro 15	
					TGT Cys											143
					TTC Phe											191
	CCG Pro															197
(2)	INFO	ORMA:	rion	FOR	SEQ	ID i	NO:	127:								
	i )	L) SI	(A) (B) (C)	LENC TYPE STRA	CHARA GTH: E: NU ANDEI OLOGY	121 JCLEI DNESS	base [C A0 S: D0	e pai CID DUBLE								
	( j	Li) N	MOLE	CULE	TYPE	E: CI	DNA									
	7)	/i) (	(A)	ORGA	SOUE NISNA C SUE	1: Ho		•	ens							
	(:	ix) l	(B) (C)	NAMI LOCA	E/KE) ATION NTIFI ER IN	N: 65	510 ION 1	06 METHO	DD: N	Von i re 4. LITI	. 4					
	(:	ki)	SEQUI	ENCE	DES	CRIP'	TION	: SE	Q ID	NO:	127	:				
ATT	TAAT	AAC '	TTAA	AAAT	TG G	CCAA	TTTT	A TT	TTTA	gaaa	AGC	TCTG	CAT	CATC	CTGTG'	r 60
TTA						u Th					n Gl				A CGA a Arg 1	109
			GAG Glu 5													121
(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	128:								
	(	i) S	(A) (B) (C)	LEN TYP STR	CHAR GTH: E: N ANDE OLOG	238 UCLE DNES	bas IC A S: D	e pa CID OUBL								

(ii) MOLECULE TYPE: CDNA

<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F) TISSUE TYPE: Brain</pre>	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 146223     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 4.4</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 128:	
AAAATATGTC TTCAGCTCTA ATCCATTATC ACTCAGATCA TTCTAACCTT TTCCCCTTGC	60
TTATCTATAA CTTTCCACTT CAACAGTGAG AAACCTGGCT TCCATATCTG TCATCCATAA	120
ATGTACGTAT TTAATTCCAG TACAC ATG TAT ACT GGT TTC AGA ATA GAA GCA  Met Tyr Thr Gly Phe Arg Ile Glu Ala  -25 -20	172
ACT TTA TTA ACT AGA GTG CAG TGC TTA TGT GCA ATT CCT TTT GCC TTT Thr Leu Leu Thr Arg Val Gln Cys Leu Cys Ala Ile Pro Phe Ala Phe -15 -5	220
AGT CTT ACA GGC ATC CGG Ser Leu Thr Gly Ile Arg 1 5	238
(2) INFORMATION FOR SEQ ID NO: 129:  (i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 419 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA	
(vi) ORIGINAL SOURCE:  (A) ORGANISM: Homo Sapiens  (F) TISSUE TYPE: Brain	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 252392     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 4.4</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 129:	
AAGOGGACCA CCTGGGTGCT GTCGTAGTTG GAGGTGGCCT GAGGAGCTCA GTTCCCTCAG	60
CGCCCGTAGT TTCGGCGGAG TCTGCGCGAT GGGCGACCCG GAAAGGCCGG GAAGCGGCCG	120

100	
GGCTGGATCA GGATGAGAGA TCATCTTCAG ACACCAACGA AAGTGAAATA AAGTCAAATG	180
AAGAGCCRST CCTAAGAAAG AGTTCTCGCC GGTTTGTCAT CTTTCCAATC CAGTACCCTG	240
ATATTTGGAA A ATG TAT AAA CAG GCA CAG GCT TCC TTC TGG ACA GCA GAA  Met Tyr Lys Gln Ala Gln Ala Ser Phe Trp Thr Ala Glu  -45 -40 -35	290
GAG GTC GAC TTA TCA AAG GAT CTC CCT CAC TGG AAC AAG CTT AAA GCA Glu Val Asp Leu Ser Lys Asp Leu Pro His Trp Asn Lys Leu Lys Ala -30 -25 -20	338
GAT GAG AAG TAC TTC ATC TCT CAC ATC TTA GCC TTT TTT GCA GCC AGT Asp Glu Lys Tyr Phe Ile Ser His Ile Leu Ala Phe Phe Ala Ala Ser -15 -10 -5	386
GAT GGA ATT GTA AAT GAA AAT TTG GTG GAG CGC Asp Gly Ile Val Asn Glu Asn Leu Val Glu Arg 1 5	419
(2) INFORMATION FOR SEQ ID NO: 130:	
<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 255 base pairs</li> <li>(B) TYPE: NUCLEIC ACID</li> <li>(C) STRANDEDNESS: DOUBLE</li> <li>(D) TOPOLOGY: LINEAR</li> </ul>	
(ii) MOLECULE TYPE: CDNA	
<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F) TISSUE TYPE: Brain</pre>	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 112195     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 4.3</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 130:	
ARTTACGATG TKKTGTGTGC TTGTGCAAAT ACAGGACGGT TCCTGAAATG TGTCTCTGAG	60
CGTTCTTAAC TGTGTGTRAG GAATTSMTGC GCGTACACGT GGTGGGTCAT T ATG CTG Met Leu	117
CTG CAC CTG TGT AGT GTG AAG AAT CTG TAC CAG AAC AGG TTT TTA GGC Leu His Leu Cys Ser Val Lys Asn Leu Tyr Gln Asn Arg Phe Leu Gly -25	165
CTG GCC GCC ATG GCG TCT CCT TCT AGA AAC TCC CAG AGC CGA CGC CGG Leu Ala Ala Met Ala Ser Pro Ser Arg Asn Ser Gln Ser Arg Arg Arg -10 -5 1 5	213
TGC AAG GAG CCG CTC CGA TAC AGC TAC AAC CCC GAC CAG GGG Cys Lys Glu Pro Lew Arg Tyr Ser Tyr Asn Pro Asp Gln Gly	255

10 15 20

(2) INFORMATION	FOR	SEQ	ID	NO:	131:
-----------------	-----	-----	----	-----	------

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 287 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 123..176
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.3

seq WTCLKSFPSPTSS/HA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 131:

AAGAGCATCC TGCGCCCCGG CGCGGGGCCC TGCGGTAGCC TCAGGCCCCT CCCCTGGACC 60

CGCCGCAGAG CCAGTGCAGA ATACAGAAAC TGCAGCCATG ACCACGCACG TCACCCTGGA 120

AG ATG CCC TGT CCA ACG TGG ACC TGC TTG AAG AGC TTC CCC TCC CCG

Met Pro Cys Pro Thr Trp Thr Cys Leu Lys Ser Phe Pro Ser Pro

-15

-10

-5

ACC AGC AGC CAT GCA TCG AGC CTC CAC CTT CCT CCA TCA TGT ACC AGG
Thr Ser Ser His Ala Ser Ser Leu His Leu Pro Pro Ser Cys Thr Arg

1 5 10

CTA ACT TTG ACA CAA ACT TTG AGG ACA GGA ATG CAT TTG TCA CGG GCA
Leu Thr Leu Thr Gln Thr Leu Arg Thr Gly Met His Leu Ser Arg Ala

20
25

TTG CAA GGT ACA TTG ACC AGG CAG
Leu Gln Gly Thr Leu Thr Arg Gln
30 35

# (2) INFORMATION FOR SEQ ID NO: 132:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 224 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:

(A)	ORGANISM: Hom	no Sapiens
(F)	TISSUE TYPE:	Brain

### (ix) FEATURE:

(A) NAME/KEY: sig peptide

(B) LOCATION: 6..104

- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 4.3

seq LLGWGLNLTLGQG/AP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 132:

AAAGG ATG GAG GAT CTC TTT AGC CCC TCA ATT AWG CCG CCG GCG CCC AAC

Met Glu Asp Leu Phe Ser Pro Ser Ile Xaa Pro Pro Ala Pro Asn

-30

-25

ATT TCC GTG CCC ATC TTG CTG GGC TGG GGT CTC AAC CTG ACC TTG GGG

Ile Ser Val Pro Ile Leu Leu Gly Trp Gly Leu Asn Leu Thr Leu Gly

-15

-10

-5

CAA GGA GCC CCT GCC TCT GGG CCG CCC AGC CGC GTC CGC CTG GTG
Gln Gly Ala Pro Ala Ser Gly Pro Pro Ser Arg Arg Val Arg Leu Val

1 5 10

TTC CTG GGG GTC ATC CTG GTG GTG GCG GTG GCA KGC AAC ACC ACA GTG

Phe Leu Gly Val Ile Leu Val Val Ala Val Ala Xaa Asn Thr Thr Val

15 20 25 30

CTG TGC CGC CTG TGC GGC GGC GGC CCG
Leu Cys Arg Leu Cys Gly Gly Gly Pro
35 40

# (2) INFORMATION FOR SEQ ID NO: 133:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 347 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 183..338
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.1

seq VMLETCGLLVSLG/HP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 133:

WO 99/06552	103	PC 1/1D70/
AGAACTGTGC TGGGAAGGAT GG	STAGGGCGA CTGGGGCTCA CCTCCGCACC GTTGTAG	GAC 120
CCGGGGTAGG GTTTTGAGCC CG	GTGGGAGCK GCCCACGCG GCCTCGTCCT GCCAACG	GTC 180
	GAC GCA GCG CAG ATG TTG GTG ACC TTC AAG Asp Ala Ala Gln Met Leu Val Thr Phe Lys -45 -40	
	ACC CGG GAG GAG TGG AGA CAG CTG GAC CT Thr Arg Glu Glu Trp Arg Gln Leu Asp Le -30 -25	
	CGA GAG GTG ATG CTG GAG ACC TGT GGG CT Arg Glu Val Met Leu Glu Thr Cys Gly Le -15	
CTG GTT TCA CTA GGG CAT Leu Val Ser Leu Gly His -5 1		347
(2) INFORMATION FOR SEQ	ID NO: 134:	
(B) TYPE: NU (C) STRANDED (D) TOPOLOGY (ii) MOLECULE TYPE (vi) ORIGINAL SOUR	432 base pairs ICLEIC ACID DNESS: DOUBLE T: LINEAR C: CDNA	
(B) LOCATION	7: sig_peptide 1: 298336 CCATION METHOD: Von Heijne matrix	
(xi) SEQUENCE DESC	CRIPTION: SEQ ID NO: 134:	
AATTGARRTG TTTGATAACT GI	CACTTTAG GGTTTCAACC AAAACCTTGA CTTTATC	ATC 60
TTGTTATACA TTTTTCAAAA TO	GAGGTTAGA GATCAGGGGA ATGAATAGGA GAGAAGT	ACA 120
TATTTCAGTT CACTGGGCAT AC	GTGAATAG AGGAAGGAGA AAATGAACAT ACCCAAT	CCA 180
CAGAGAAATG GCTCACAGAG CC	CCAGTGACT ATGCTGAGAC GCTATTAATT CAAGAAA	GTT 240
TTAGTATTTG ATTTGTCAAA TO	GACATTATT GTTTAGGACT TTTATTTTCC CTTACAG	297
	AAT ATT GCC CAA CTG GAG GCC CAG GTG GA Asn Ile Ala Gln Leu Glu Ala Gln Val Gl	

AAG GTT ACA ANG GAA AAG ATT TCA GCT ATT AAT CAA CTG GAG GAA AAT 393

T	WO 99	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,	) <u>L</u>						104	4					PCT	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,
гур	Val 5	Thr	Lys	Glu	Lys	Ile 10	Ser	Ala			Gln 15	Leu	Glu	Glu	Asn	
				GGC Gly												432
(2)	INFC	RMAI	ON	FOR	SEQ	ID N	10: 1	.35:								
	(i	) SE	(A) (B) (C)	ICE C LENG TYPE STRA TOPC	TH: : NU .NDED	380 CLEI NESS	base C AC : DO	pai ID UBLE								
	(i	i) M	OLEC	ULE	TYPE	: CE	ANG									
	( v	i) C	(A)	NAL ORGA TISS	NISM	: Ho		-	ns							
		ж) F	(A) (B) (C) (D)	NAME LOCA IDEN OTHE	TION TIFI R IN	: 90 CATI	O15 ON M	2 IETHC ON:	D: V scor seq	e 4. GLWA	1 HSWT	cscs				
	(х	1) 8	SEQUE	ENCE	DESC	RIPI	TON:	SEC	) ID	NO:	135:					
AATI	CACI	TC P	ACCT(	GGAGT	TT GA	AGCC	AAGAT	TCT	CTT	TACT	CCA	AAGCC	AG C	CACTO	CCTTCT	60
GAGA	ACAC	GA C		SATT	rg ga	ATGG	ACGG	ATG	ተጥር	CTG	ccc	GCC				
								Met				Ala	Ser	-15	Gln	113
Gly	Leu	Trp	GCT Ala	CAC His	Ser	Trp	Thr	TGC Cys	Leu -20 AGC Ser	Leu TGT Cys	Gly TCA Ser	GCG Ala	GCA Ala	-15 STG Xaa	CGG	113
Gly	Leu	Trp	GCT Ala -10 CCA	His	Ser GGA	Trp	Thr	TGC Cys -5 ATG	Leu -20 AGC Ser	TGT Cys	TCA Ser	GCG Ala CAG	GCA Ala 1 GCG	-15 STG Xaa GGG	CGG Arg	
TCT Ser	GTC Val 5 CCT	Trp CAC His	GCT Ala -10 CCA Pro	His GGC	Ser GGA Gly CCT	GAC Asp 10	Thr TGG Trp	TGC Cys -5 ATG Met	Leu -20 AGC Ser CAA Gln	TGT Cys CAG Gln	TCA Ser TTT Phe 15	GCG Ala CAG Gln	GCA Ala 1 GCG Ala	-15 STG Xaa GGG Gly	CGG Arg TTC Phe	161
TCT Ser CTC Leu 20	GTC Val 5 CCT Pro	CAC His CCC Pro	GCT Ala -10 CCA Pro CAG Gln	His GGC Gly GTG	GGA Gly CCT Pro 25 CTG	GAC Asp 10 GCC Ala	Thr TGG Trp CAC His	TGC Cys -5 ATG Met CTC Leu	Leu -20 AGC Ser CAA Gln TCC Ser	TGT Cys CAG Gln CTT Leu 30 CTT Leu	TCA Ser TTT Phe 15 ACA Thr	GCG Ala CAG Gln TGG Trp	GCA Ala 1 GCG Ala GAT Asp	-15 STG Xaa GGG Gly GTC Val	CGG Arg TTC Phe TCT Ser 35	161 209
TCT Ser CTC Leu 20 CTT Leu	GTC Val 5 CCT Pro CTT Leu	CAC His CCC Pro	GCT Ala -10 CCA Pro CAG Gln CCT Pro	GGC Gly GTG Val TGC Cys 40 GAT Asp	GGA Gly CCT Pro 25 CTG Leu	GAC Asp 10 GCC Ala GTC Val	Thr TGG Trp CAC His CCT Pro	TGC Cys -5 ATG Met CTC Leu AAA Lys	Leu -20 AGC Ser CAA Gln TCC Ser GCA Ala 45 ACC Thr	TGT Cys CAG Gln CTT Leu 30 CTT Leu	TCA Ser TTT Phe 15 ACA Thr GAG Glu	GCG Ala CAG Gln TGG Trp TTT Phe	GCA Ala 1 GCG Ala GAT Asp GTG Val	-15 STG Xaa GGG Gly GTC Val GTT Val 50	CGG Arg TTC Phe TCT Ser 35 CAT His	161 209 257

380

75 70

(2)	INFO	ORMAT	CION	POR	SEQ	ID t	NO: 1	136:							
	(i	.) SE	(A) (B) (C)	LENG TYPE STRA	TH: : NU NDEC	212 CLEI NESS	RISTI base [C AC S: DC [NEAF	pai ID UBLE							
	( i	.i) M	OLEC	ULE	TYPE	: CI	ANC								
	(1	/i) C	(A)	ORGA	NISM	i: Hc	omo S Bra	•	ens						
		(i) S	(A) (B) (C) (D)	NAME LOCA IDEN OTHE	TION TIFI R IN	: 9. CATI	M NOI	IETHO ON:	D: V scor seq	e 4. APLE	l CLSCW	IGGGW			
AGC(	GCAA		: Ala					ı Lei					/ Gl	C TGG / Trp	
	CTC Leu 1														98
	TTT Phe														146
	GGT Gly														194
	TCT Ser														212
(2)	INF	ORMA'	TION	FOR	SEQ	ID	NO:	137:							
	(,	i) S	EQUEI	NCE (	CHAR	ACTE	RIST	ICS:							

- - (A) LENGTH: 432 base pairs
    (B) TYPE: NUCLEIC ACID
    (C) STRANDEDNESS: DOUBLE
    (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain

# (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(3) LOCATION: 226..285

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.1

seq LGFLNCYIAVARS/GG

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 137:

AAGGGAMN	ISA (	CCCAC	GCT	SC GC	GACS	SGGT	CAC	GCT	GCGG	CGC	rgaco	GGC (	CTCTC	CTCCT	60
TCCGCGGG	STT 1	CCG	ACTCO	CC TC	GCCC1	TAGAT	TTT	CTG	CTTA	GCG	ACTTO	GG (	STCC	CTCTC	120
GTTTGCT1	CT (	GGTAC	GAG	rc Go	CAATO	CCA	K BAC	CAA	ragc	CCA	GAAGA	AGG A	ACACO	GTTCC	180
CGTACCG	AAG (	GTTC <i>I</i>	AGTAC	CC AC	GCAGO	CCG	A CC	ATCAC	CGCG	GCG	Me			OR GTT na Val	
GGC ATT Gly Ile -15															285
GGC GGC Gly Gly 1															333
GCC TGT Ala Cys															381
AAG AGC Lys Ser															429
AAG Lys															432

# (2) INFORMATION FOR SEQ ID NO: 138:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 229 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) CRIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide
(B) LOCATION: 101..157

107

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.1

seq FVVFSTMFTASSP/GE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 138:

AGATGAATAT TTGATACCCA CAGTAGAACT TCTTTMMGAA CTTCTTTCAC AATWGGAGTT 60 AATCTTCTAA AGCCCCGCCA CTGCTTCATC AACTAAGTTT ATG GAA TAT TCT AAA Met Glu Tyr Ser Lys TMM TTT GTT GTC TTT TCA ACA ATG TTC ACA GCA TCT TCA CCA GGA GAA 163 Xaa Phe Val Val Phe Ser Thr Met Phe Thr Ala Ser Ser Pro Gly Glu -10 -5 GAC TTT CCC CCC TTC TTT TCA CAG ATG TNS AGA TTG TCA AGA AAC TAC Asp Phe Pro Pro Phe Phe Ser Gln Met Xaa Arg Leu Ser Arg Asn Tyr 10 TTT CCT TGC CCA CCR WGG 229

(2) INFORMATION FOR SEQ ID NO: 139:

Phe Pro Cys Pro Pro Xaa

20

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 328 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 113..232

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.1

seq LPFRLPWASTATA/RC

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 139:

AACACCCAGG CCCTGATGCA GCAGCAGGCG GCCCTGGTAG CGGCTCACAG TGCCTACCTC

AGCCCCATGG CCACCATGGC TGCCGTGCAG ATGCAGCACA TGGCTGCCAT CA ATG CCA 118 Met Pro -40

ATG GCC TOA TOG CCA CCC CCA TCA CCC CAT CCT CAG GAA CCA GCA CCC 166 Met Ala Ser Ser Pro Pro Pro Ser Pro His Pro Gln Glu Pro Ala Pro -35 -30

WO 99/06552	108	PCT/IB98/01236
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CTC	CTG	CCA	TCG	CTG	CCA	CGC	CTG	TCT	CTG	CCA	TTC	CGG	CTG	CCC	TGG	214
Leu	Leu		Ser	Leu	Pro	Arg		Ser	Leu	Pro	Phe	Arg	Leu	Pro	Trp	
		-20					-15					-10				
GCG	TCA	ACG	GCT	ACA	GCC	CGG	TGC	CCA	CCC	AGC	CCA	CTG	GGC	AGC	CTG	262
Ala	Ser	Thr	Ala	Thr	Ala	Arg	Cys	Pro	Pro	Ser	Pro	Leu	Gly	Ser	Leu	
	-5					i				5			-		10	
CNC	CTG	ATG	CTC	TGT	ATC	CCA	ACG	GGG	TTC	ACC	CCT	ACC	CAG	CCC	AGA	310
Xaa	Leu	Met	Leu	Cys	Ile	Pro	Thr	Gly	Phe	Thr	Pro	Thr	Gln	Pro	Arg	
				15					20					25		
ccc	ccc	CGG	CCC	ССТ	GGG											328
				Pro												320
		9	30		<u>J</u>											
			3.0													

# (2) INFORMATION FOR SEQ ID NO: 140:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 217 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide
(B) LOCATION: 53..166

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4

seq WALGLKFLSSSSQ/NF

(Mi) SEQUENCE DESCRIPTION: SEQ ID NO: 140:

AACAGGAGAC :	TTGGGAAGGA CCAA1	GGTAA TTTAAGTGGC	TCTTAAAAAG TC ATG CAA 58 Met Gln
		Asp Pro Ile Ala	ATA ATG TAT GTG TGC 106 Ile Met Tyr Val Cys -25
			CTG AAA TTC CTG AGT 154 Leu Lys Phe Leu Ser -5
			TTC CTC ATT TTA CAT 202 Phe Leu Ile Leu His 10
ACA GGA GGC Thr Gly Gly	Gln Arg		217

(2)	INFO	ORMA:	LION	FOR	SEQ	ID 1	10: 1	141:							
	i)	L) SI	(A) (B) (C)	NCE C LENC TYPE STRA TOPC	TH: : NU ANDEC	202 CLEI NESS	base C AC	pai ID UBLE							
	( i	i) N	OLE	CULE	TYPE	: C	ONA								
	(1	/i) (	(A)	NAL ORGA TISS	NISM	l: Ho		•	ens						
	(i	Lx) I	(B) (C)	JRE: NAME LOCA IDEN OTHE	TION TIFI	: 44 CATI	13 ON M	3 ETHC	D: V	e 4	leijr LKCLI				
	()	ci) S	SEQUE	ENCE	DESC	RIPI	ION:	SEC	) ID	NO:	141:	:			
ATA	\TAG1	rgt :	ratt:	rcag:	rg c <i>i</i>	ATGAT	TTTT	r GCC	CTTTC	GAGA	GAC		GGT Gly		55
				AAG Lys											103
				CTG Leu											151
				CTT Leu										GAA Glu	199
TGG Trp															202
(2)				FOR	_										
	(:	i) S	(A) (B) (C)	NCE ( LENG TYPI STRI	GTH: E: NO ANDEO	361 JCLE: ONES:	base IC AG S: DG	e pai CID DUBLE							
	(:	ii)	MOLE	CULE	TYP	E: C	DNA								
	(:	vi)	(A)	INAL ORG TIS	ANIS	4: H			ens						

U 99/00332	110	. 0 2/12/3/3/3
(ix) FEATURE:		

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 248..355
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 4

seq VQLSFAATTPVLA/DK

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 142:

AAGTTGGGAG AGGAGCTGCT GGTTGAAGTG AAGGTGAGGA GCTCAGTGCT TCTTTCACTG 60

CCCTATCTGC TGGGCTTTAC GCCCCTGAGG GGCTGACTGT AAAAAAACTCT AAGCTGATCC 120

AGCCCCCAAA ATTCACCTTT GGTGAGCTGG AAAGTCCATC TATTTGGGAC GCGAATCATG 180

TCAGTGCGAC AACGCAAAAG GGTTGAAAGC CTTCTACGAT GCAATAAAAT ACGGGCCTAA 240

CCACTTG ATG GTG TTT GGA GGC GTC TGT CCA TCC GTC ACA TCC ATC ATT 289

Met Val Phe Gly Gly Val Cys Pro Ser Val Thr Ser Ile Ile -35 -25

GCA GAG TCC CTC CAA GGC TGG AAT CTG GTG CAG CTT TCT TTT GCT GCA 337

Ala Glu Ser Leu Gln Gly Trp Asn Leu Val Gln Leu Ser Phe Ala Ala -20 -15 -10

ACC ACG CCT GTT CTA GCC GAT AAG 361

Thr Thr Pro Val Leu Ala Asp Lys -5

#### (2) INFORMATION FOR SEQ ID NO: 143:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 216 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 145..192
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4

seq ITWSLLFLYQCSL/HF

(Mi) SEQUENCE DESCRIPTION: SEQ ID NO: 143:

ACACATTACC TCTTTCATT TTAGACAGGT TAATTAGTGT GTATTTCCAT AGTTGTCTTT 60

TACCTCAAGA AATAATCATT TCTTTAGGTA ATTATTTTAA TGGCTTGCCA TTTTGTATGA 120

TTGTTCTTGC AAACATTTCT ATTT ATG CAT TTT ATA ACA TGG AGC TTA CTA 171

Met His Phe Ile Thr Trp Ser Leu Leu

TTT TTA TAC CAG TGC TCG CTT CAT TTT ATC AAT ATC AAG GCC GGG
Phe Leu Tyr Gln Cys Ser Leu His Phe Ile Ile Lys Ala Gly
-5
1
5

#### (2) INFORMATION FOR SEQ ID NO: 144:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 378 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 256..363
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.9

seq CWPSVASPSSSWS/SP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 144:

AAAGTGGCCA GCGGACCATC TCTCGTGCCC TCGCTCTCTG CGCTCCGGGG CAGCTGAGCC 60

CCGGCCACCC GCTCTCCAAG ATGAAGAAGC TCCAGGGAGC TCACCTCCGC AAGCCTGTCA 120

CCCCAGACCT GCTGATGACC CCCAGTGACC AGGGCGATGT CGACCTGGAT GTGGACTTTG 180

CTGCACACCG GGGGAACTGG ACAGGCAAGC TGGACTTCCT GCTGTCCTGC ATTGGCTACT 240

GTGTAGGCCT GGGGA ATG TCT GGC GCT TCC CCT ATC GAG CGT ACA CCA ATG 291

Met Ser Gly Ala Ser Pro Ile Glu Arg Thr Pro Met -35 -30 -25

GAG GAG GCG CCT TCC TCG TGC CCT ACT TCC TCA TGC TGG CCA TCT GTG Glu Glu Ala Pro Ser Ser Cys Pro Thr Ser Ser Cys Trp Pro Ser Val

GCA TCC CCC TCT TCT TCC TGG AGC TCT CCC TGG GCC AGT

Ala Ser Pro Ser Ser Trp Ser Ser Pro Trp Ala Ser

# (2) INFORMATION FOR SEQ ID NO: 145:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 321 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(D) TOPOLOGY: LINEAR

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide
(B) LOCATION: 172..282

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 3.9

seq PGPSLRLFSGSQA/SV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 145:

AAGGGTCTCC ATGACAACCG GCCTGGCCGG CTAGCAGTGC TCTGCTCACT TGGCTGCGAG 60

GAGCGCCACG AAAGGTCAGA GGAAGGAGCT GTGGGAAGCT CGCAGCAGGT ATCGGAGCTT 120

AAGCCAGTGG ATTTGGGGGC CCTGGGCTCC CTAGCCGGCT GCGGTGTGAG A ATG GAG Met Glu

TGG GCA GGA AAG CAG CGG GAC TTT CAG GTA AGG GCA GCT CCG GGC TGG

Trp Ala Gly Lys Gln Arg Asp Phe Gln Val Arg Ala Ala Pro Gly Trp -35 -20

GAT CAT TTG GCC TCC TTT CCT GGC CCT TCT CTC CGG CTG TTT TCT GGG Asp His Leu Ala Ser Phe Pro Gly Pro Ser Leu Arg Leu Phe Ser Gly -15 -5

AGT CAG GCG AGT GTC TGT AGT CTC TGC TCG GGG TTT GGG GCT CAG GAA 321

Ser Gln Ala Ser Val Cys Ser Leu Cys Ser Gly Phe Gly Ala Gln Glu

#### (2) INFORMATION FOR SEQ ID NO: 146:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 278 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 78..257
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.9

seq AKVVSLSLQTSSA/HH

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 146:

AAGAACACAA AAAGCTACAG AAGGTCCAGG CTACTGAAAA GCATCAAGAC CAAGCTGTTG	60
TAAGTCAAAC TGCTTTT ATG ATT GCA TTC TTT GAT GAA GAC AAT CCC AGA Met Ile Ala Phe Phe Asp Glu Asp Asn Pro Arg -60 -55 -50	110
AAA AGA AGG TCG TAT TCT TTT ACT CAA AGT GCG GGA ATC TTG TGT CAG Lys Arg Arg Ser Tyr Ser Phe Thr Gln Ser Ala Gly Ile Leu Cys Gln -45 -35	158
GAA ACT ACA TAT TCA ACA CCA CAT ACA AAA CTT GAG AAA GCA AAG TCT Glu Thr Thr Tyr Ser Thr Pro His Thr Lys Leu Glu Lys Ala Lys Ser -30 -25 -20	206
CCA ACA GCA GAT GCC AAA GTG GTT TCT TTG TCT TTA CAG ACT AGC TCT Pro Thr Ala Asp Ala Lys Val Val Ser Leu Ser Leu Gln Thr Ser Ser -15 -10 -5	254
GCG CAT CAC AGA GGG GGG MDT GGT Ala His His Arg Gly Gly Xaa Gly 1 5	278
(2) INFORMATION FOR SEQ ID NO: 147:  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 349 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain  (ix) FEATURE: (A) NAME/KEY: sig_peptide (B) LOCATION: 89232 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 3.9 seq ALFCTLPCPVERG/QQ  (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 147:	
AAATTTTCTT AAATTAGCAG TCTAATGTGT TCTAAAGAGC AGCTCCACTC AGATCTCTTC	60
TTAAGGGTTG CATTTCATCA CAGTATAA ATG GGC AAA TCC ATC AYT TCC CTC Met Gly Lys Ser Ile Xaa Ser Leu -45	112
TGC TCA GTG CKC TTG AAR GCG AGA TTG AAG GGM MAA TTA GAA GCT GTT Cys Ser Val Xaa Leu Lys Ala Arg Leu Lys Gly Xaa Leu Glu Ala Val -40 -35 -30 -25	160
CAT TTG TGC TTG CGG GCT CAG AAG CGT CGC ACT GCT TTG TTT TGT ACT	203

His	Leu	Cys	Leu	Arg -20	Ala	Gln	Lys	Arg	Arg -15	Thr	Ala	Leu	Phe	Cys -10	Thr	
CTA Leu	CCG Pro	TGT Cys	CCT Pro -5	GTT Val	GAA Glu	AGG Arg	GGT Gly	CAA Gln l	CAA Gln	GTG Val	CCG Pro	GGG Gly 5	ANV Xaa	NNN Xaa	AHG Xaa	256
AGG Arg	CTG Leu 10	AGG Arg	CTG Leu	GCG Ala	TCA Ser	CCT Pro 15	TCC Ser	GTT Val	GCT Ala	AAG Lys	GTG Val 20	TTC Phe	CAG Gln	TGT Cys	TTT Phe	304
							AAC Asn									349
(2)	INFO	RMAT	CION	FOR	SEQ	ID N	NO: 1	148:								
	(i	.) SE	(A) (B) (C)	LENG TYPE STRA	TH: : NU NDEC	210 CLEI NESS	Dase C AC C DC	pai ID UBLE								
	(i	.i) M	OLEC	ULE	TYPE	e: co	NA									
	( v	7i) C	(A)	ORGA	NISM	1: Hc	omo S Bra	•	ens							
	(i	.x) F	(A) (B) (C)	NAME LOCA I DEN	TION TIFI	: 52 CATI	.g_pe 96 ON M MATIC	IETHC	D: V	e 3.	9		itrix S/XE			
	()	(i) S	EQUE	CNCE	DESC	CRIPT	:NOI	SEÇ	) ID	NO:	148:	:				
ACAC	CAAAC	GGA #	\AAT(	GCA	GG GI	TATE	SAATI	r TC	гттс	CCAG	CTT	TAT	ATA (		G TGC t Cys 5	57
CTG Leu	CAT His	ATG Met	ACA Thr -10	CTC Leu	TTT Phe	AGA Arg	GTT Val	CCT Pro -5	TTC Phe	ACT Thr	TTT Phe	TCT Ser	KTT Xaa 1	TTT Phe	TGG Trp	105
							GAG Glu									153
TAC Tyr 20	TAC Tyr	AAA Lys	CTT Leu	ATT Ile	ATG Met 25	GTA Val	CTT Leu	AAA Lys	ATT Ile	GCA Ala 30	CTC Leu	CTC Leu	CTG Leu	TCC Ser	CCG Pro 35	201
	CCC Pro						•									210

(2) INFORMATION FOR SEQ ID NO: 149:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 143 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 75..116

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 3.9

seq LNILKTLTSAALP/SP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 149:

AATACACCTT AAACTTTACT ACTTTTTATA AACGGTAGGA AAGGATATAC TGATGTTGTG 60

GGTATTACAA GGTA ATG CTG AAC ATT CTG AAG ACC TTA ACT TCT GCT GCT 110

Met Leu Asn Ile Leu Lys Thr Leu Thr Ser Ala Ala

-10 -

CTT CCC TCC CCC TCC CCC CGC CCC AAC AAG AGG
Leu Pro Ser Pro Ser Pro Arg Pro Asn Lys Arg

1 5

(2) INFORMATION FOR SEQ ID NO: 150:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 176 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig peptide

(B) LOCATION: 24..143

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 3.9

seq SPLLCLYHPPVYT/ST

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 150:

110	
AGTAATCCCA GGCGTTCGCC CTC ATG CGG GCC AGG GTT TGG CCT CGC TCC CAC  Met Arg Ala Arg Val Trp Pro Arg Ser His  -40  -35	53
GGG ATC CCT GTG CCT TCC TTT CTC TCT AAG AGC AGC CTC AGT CAT ACA Gly Ile Pro Val Pro Ser Phe Leu Ser Lys Ser Ser Leu Ser His Thr -30 -25 -20 -15	101
CCA TCA CCT CTC CTC TGT CTA TAC CAT CCT CCT GTC TAC ACC AGC ACC Pro Ser Pro Leu Cys Leu Tyr His Pro Pro Val Tyr Thr Ser Thr -10 -5 1	149
ACT ACC CCA TCT ATA CCA CCA CGT CTG Thr Thr Pro Ser Ile Pro Pro Arg Leu 5 10	176
(2) INFORMATION FOR SEQ ID NO: 151:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 414 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: CDNA	
<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F) TISSUE TYPE: Brain</pre>	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 262369     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 3.9</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 151:	
AAAGTAGGAA ATGGCTGCTT CACCCAGGAG GCACCAAGAT GCCCGTGTGT GGCTCTACTG	60
GTGATGCCCT GGTCTTCATT GAAAAGGCCA GCACCCGTTA CGTGGTCAGC ACAGACGTTG	120
CCGTGAATGA GGATTCCTTC CTACAGATAG ACTTCGCTGC CTCCTGCTCA GTCACAGACT	180
CTTGTTATGC GATTGAATTG GAATACTCAG TAGATCTTGG ATTGTCATGG CACCCATTGG	240
TAAGGGACTG TCTGCCTACC A ATG TGG AAT GCA GTC GCT ATC ATC TGC AAC  Met Trp Asn Ala Val Ala Ile Ile Cys Asn  -35  -30	291
GGA TCC TGG TGT CAG ACA CDW TCA ACA AGT GGA CTA GAA TCA CTC TGC Gly Ser Trp Cys Gln Thr Xaa Ser Thr Ser Gly Leu Glu Ser Leu Cys -25 -15	339
CTC TCC CTC CTT ATA CCA GGT CCC AAG CCA CTC GTT TCC GTT GGC ATC Leu Ser Leu Leu Ile Pro Gly Pro Lys Pro Leu Val Ser Val Gly Ile -10 -5 1 5	387

WO 99/06552

AAC CAG CTC CTT TTG ACA AGC AGC AGA Asn Gln Leu Leu Thr Ser Ser Arg 10 15	414
(2) INFORMATION FOR SEQ ID NO: 152:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 171 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: CDNA	
<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F) TISSUE TYPE: Brain</pre>	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 103144     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 3.9</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 152:	
AAAAATTATT TCTGCTTAAA CAACAGTTTC AAATATTTCT CTTTTGAAGA CAAAATTGGT	60
TTAGTTTCAG CAATGTATTG ATATAATTTT ACATTTTTTT AA ATG TTG AGG CTG Met Leu Arg Leu	114
GGT TTA TTT AAG ATT AGC TGG GCT CGC TGC CTA TCA TAT AGT AAA ACC Gly Leu Phe Lys Ile Ser Trp Ala Arg Cys Leu Ser Tyr Ser Lys Thr -10 -5 5	162
CAG CBC GAA Gln Xaa Glu	171
(2) INFORMATION FOR SEQ ID NO: 153:	
<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 262 base pairs</li><li>(B) TYPE: NUCLEIC ACID</li><li>(C) STRANDEDNESS: DOUBLE</li><li>(D) TOPOLOGY: LINEAR</li></ul>	
(ii) MOLECULE TYPE: CDNA	
<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F) TISSUE TYPE: Brain</pre>	

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 80..187

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 3.9

seq VVEILPYLPCLTA/RD

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 153:

AGGGGTACCG AGTCTCGTTT CCTCTCAGTC CATCCACCCT TCATGGGGCC AGAGCCCTCT 60 CTCCAGAATC TGAGCAGCA ATG CCG TTT GCT GAA GAC AAG ACC TAT AAG TAT 112 Met Pro Phe Ala Glu Asp Lys Thr Tyr Lys Tyr -35 -30 ATC TGC CGC AAT TTC AGC AAT TTT TGC AAT GTG GAT GTT GTA GAG ATT 160 Ile Cys Arg Asn Phe Ser Asn Phe Cys Asn Val Asp Val Val Glu Ile -20 -15 CTG CCT TAC CTG CCC TGC CTC ACA GCA AGA GAC CAG GAT CGA CTG CGG 208 Leu Pro Tyr Leu Pro Cys Leu Thr Ala Arg Asp Gln Asp Arg Leu Arg GCC ACC TGC ACA CTC TCA GGG AAC CGG GAC ACC CTC TGG CAT CTC TTC 256 Ala Thr Cys Thr Leu Ser Gly Asn Arg Asp Thr Leu Trp His Leu Phe 10 AAT ACC 262 Asn Thr

#### (2) INFORMATION FOR SEQ ID NO: 154:

25

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 165 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 46..153
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.8

seq GTDSLSFLPPCPC/CP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 154:

AATCATGAAA TCCTTGCAAC TCATTAAGTT TCCTGTTTGC TGTAG ATG CCA GGA AGC 57

Met Pro Gly Ser

-35

TCA GGG CTC AGA TTT ATA TGT AAG TCC AGG AAC CAT CCT CAG TTT GGG 105

Ser Gly Leu Arg Phe Ile Cys Lys Ser Arg Asn His Pro Gln Phe Gly
-30 -25 -20

AGT TTC AGT GGA ACT GAC TCC CTT TCC TTC CTA CCA CCC TGC CCC TGT

Ser Phe Ser Gly Thr Asp Ser Leu Ser Phe Leu Pro Pro Cys Pro Cys

-15

-10

-5

10 10

TGC CCG GCT GCG

Cys Pro Ala Ala

1

## (2) INFORMATION FOR SEQ ID NO: 155:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 261 base pairs
- (B) TYPE: NUCLEIC ACID
- (C) STRANDEDNESS: DOUBLE
- (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 64..234
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.8

seq QLXLVMEFCGAGS/VT

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 155:

ACATACGGGC AAGTTTATAA GGGTCGTCAT GTCAAAACGG GCCAGCTTGC AGCCATCAAG 60

GTT ATG GAT GTC ACA GGG GAT GAA GAG GAA GAA ATC AAA CAA GAA ATT

Met Asp Val Thr Gly Asp Glu Glu Glu Ile Lys Gln Glu Ile

-55

-50

-45

AAC ATG TTG AAG AAA TAT TCT CAT CAC CGG AAT ATT GCT ACA TAC TAT

Asn Met Leu Lys Lys Tyr Ser His His Arg Asn Ile Ala Thr Tyr Tyr

-40

-30

GGT GCT TTT ATC AAA AAG AAC CCA CCA GGC ATG GAT GAC CAA CTT TGR 204
Gly Ala Phe Ile Lys Lys Asn Pro Pro Gly Met Asp Asp Gln Leu Xaa
-25 -20 -15

TTG GTG ATG GAG TTT TGT GGT GCT GGC TCT GTC ACC GAC CTG ATC AAG

Leu Val Met Glu Phe Cys Gly Ala Gly Ser Val Thr Asp Leu Ile Lys

-10

-5

1

5

AAC ACA GGG 261 Asn Thr Gly

(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 126 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR (ii) MOLECULE TYPE: CDNA (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain (ix) FEATURE: (A) NAME/KEY: sig\_peptide (B) LOCATION: 49..120 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 3.8 seg KLFLVFLLNICKG/IV (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 156: ATCTCTAGAA AGAAGAAGGC ATGCTACAAA TAGGAAGGAA TTGTAATA ATG ATA TTT 57 Met Ile Phe GGC CTC TAC TTT GTC TTA GCT GTT AAA CTG TTT TTA GTA TTT TTG TTA 105 Gly Leu Tyr Phe Val Leu Ala Val Lys Leu Phe Leu Val Phe Leu Leu AAT ATT TGC AAA GGG ATC GTG 126 Asn Ile Cys Lys Gly Ile Val -5 1 (2) INFORMATION FOR SEQ ID NO: 157: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 383 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR (ii) MOLECULE TYPE: CDNA (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain (ix) FEATURE: (A) NAME/KEY: sig\_peptide (B) LOCATION: 246..347 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 3.6

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 157:

seq IKCSSWISSLASG/IP

TTCAGATCTT GGGTTCAGGA GCGAGACCCA TGTCTGAAAT ATAAACTTGC TCACTGTCAG	120
CCTGTGATGG TCTTTGTGAG ACATAGAATG AATATTAATA AAGAGGTGTA AGGACTGATC	180
CTGGGATCAT CCACAGTAAG GCTGGGGGAA GAGGAGACCT GGCAAAGGAA TCAAAGACAT	240
GATCC ATG AGG AAG AAG CGA GTR GAA GAA CTA ATA GTG TTT CCA GGA GAA Met Arg Lys Lys Arg Val Glu Glu Leu Ile Val Phe Pro Gly Glu -30 -25 -20	290
GTA ACT TCT TCC TCC ATC AAG TGC TCC TCT TGG ATT TCT TCC CTG Val Thr Ser Phe Ser Ser Ile Lys Cys Ser Ser Trp Ile Ser Ser Leu -15 -10 -5	338
GCT TCT GGA ATA CCA CAC TCT CTT GGA TTC TCC CTT CCC CCA GGG Ala Ser Gly Ile Pro His Ser Leu Gly Phe Ser Leu Pro Pro Gly 1 5 10	383
(2) INFORMATION FOR SEQ ID NO: 158:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 427 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE:  (A) ORGANISM: Homo Sapiens  (F) TISSUE TYPE: Brain  (ix) FEATURE:  (A) NAME/KEY: sig_peptide  (B) LOCATION: 257340  (C) IDENTIFICATION METHOD: Von Heijne matrix  (D) OTHER INFORMATION: score 3.6  seq ACLFSXFLAVSRH/PN  (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 158:	
AAGAACCCTT TTTATAGATA GGTCTTGTCT GGATTTGTGC ACGTGGATTT ATAATGAGAG	60
ATTTTCTAGT TGTTTTTGGT TCTCCTCCTC CTCCTCCTC TTTDHCCTCC TTCTVTTCCT	120
CCTTTTCTTC CTCCTTWTCT TCTAAAACCT CTAATCTCTT ATTCCCTCTA ATGTCTGACC	180
AAAGTACTGC TGTCTGAGAC ATTGGAGGCA TACTGTGCTC CTCTTCTTCC CTCCCTGTGG	240
AGAAGCCTTA AGTTAT ATG CCT TCA TCC AGT CTT GCA GAG TTG TGT CTA ATG Met Pro Ser Ser Leu Ala Glu Leu Cys Leu Met -25 -20	292
CAG CAA GAT GCC TGC CTG TTT TCT KTG TTC CTA GCW GTC TCC AGG CAT Gln Gln Asp Ala Cys Leu Phe Ser Xaa Phe Leu Ala Val Ser Arg His -15 -5	340

	AAC Asn														388
	GTT Val														427
(2)	INFO	RMAT	ON	FOR	SEQ	ID 1	10: 1	159:							
	(i	) SE	(A) (B) (C)	LENC TYPE STRA	CHARA GTH: C: NU ANDED OLOGY	158 ICLEI NESS	base C AC S: DC	e pai CID DUBLE							
	(i	i) M	OLEC	CULE	TYPE	: CI	ANC								
	( v	i) C	(A)	ORGA	SOUF NISM SUE T	1: Hc		-	ens						
	(i	ж) <b>Е</b>	(A) (B) (C)	NAME LOCA IDEN	E/KEY ATION NTIFI ER IN	1: 2] [CAT]	14 ION N	10 Vetho	D: / scoi	re 3.	-				
	(×	i) S	EQUE	ENCE	DESC	CRIP:	CION	: SE	O ID	NO:	159	:			
AAT'	TTTCA	GT F	AGGA	AACA'	Me					er C				TG ATG al Met -30	
	ATG Met									Ser				Ala	101
	CAG Gln								Leu				Glu		149
	CTT Leu 5														158
(2)	INFO	ORMA'	TION	FOR	SEQ	ID	NO:	160:							
	( i	L) Si	(A) (B) (C)	LEN TYP STR	CHAR GTH: E: N ANDE OLOG	319 UCLE DNES	bas IC A S: D	e pa CID OUBL							

(ii) MOLECULE TYPE: CDNA

(VI) ORIGINAL SOURCE:  (A) ORGANISM: Homo Sapiens  (F) TISSUE TYPE: Brain	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 209289     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 3.6</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 160:	
AAGTTCTGTG GGCTCTATTC GGCCATATTA ATAAAGAGAA AGGGAAGGCT GACHGTCC	TT 60
CGCCTCCGCC CCCACATACA CACCCCTTCT TCCCACTCCG CTCTCACGAC TAAGCTCTC	CA 120
CGATTAAGGC ACGCCTGCCT CGATTGTCCA GCCTCTGCCA GAAGAAAGCT TAGCAGCC	AG 180
CGCCTCAGTA GAGACCTAAG GGCGCTGA ATG AGT GGG AAA GGG AAA TGC CGA Met Ser Gly Lys Gly Lys Cys Arg -25 -20	232
CCA ATT GCG CTG CGG CGG GCT GTG CCA TTA CCT ACA ACA AGC ACA TTA Pro Ile Ala Leu Arg Arg Ala Val Pro Leu Pro Thr Thr Ser Thr Leu -15 -10 -5	280
ACA TCA GCT TCC ACA GGT TTC CTT TGG ATC CTA AAA GAA Thr Ser Ala Ser Thr Gly Phe Leu Trp Ile Leu Lys Glu 1 5 10	319
(2) INFORMATION FOR SEQ ID NO: 161:  (i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 91 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA	
<ul><li>(vi) ORIGINAL SOURCE:</li><li>(A) ORGANISM: Homo Sapiens</li><li>(F) TISSUE TYPE: Brain</li></ul>	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 1467     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 3.6</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 161:	
AGAAAATGGA AAA ATG ACC CCA AAG GCA ATT CAG AAA TCA TCA GGG CTC Met Thr Pro Lys Ala Ile Gln Lys Ser Ser Gly Leu	49

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-15 -10

TTC TGC CCA TCA CAG GCC CAG AGC GCA AGA CCC GCA GAA AAG

Phe Cys Pro Ser Gln Ala Gln Ser Ala Arg Pro Ala Glu Lys

-5

1

5

## (2) INFORMATION FOR SEQ ID NO: 162:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 364 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:

Glu Val Ser His Ala Gly

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 56..271
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 3.6

seq CTSLLQLYDASNS/EW

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 162:

AAACTCGCTG GGCTCCAAAA GAAACACTGG CTTCTCTCCT	TCAGCTCCAG GCTAG ATG 58 Met
CCA GAC CAG TTT GAC CAG GCG GTT GTG CTG AAC Pro Asp Gln Phe Asp Gln Ala Val Val Leu Asn -70 -65	
GGG ATG CTG GAG ACT GTG AGA ATC CGC AAA GCT Gly Met Leu Glu Thr Val Arg Ile Arg Lys Ala -55 -50 -45	Gly Tyr Ala Val Arg
AGA CCC TTT CAG GAC TTT TAC AAA AGG TAT AAA Arg Pro Phe Gln Asp Phe Tyr Lys Arg Tyr Lys -35 -30	
CTG GCT CTG CCT GAG GAC GTC CGA GGG AAG TGC Leu Ala Leu Pro Glu Asp Val Arg Gly Lys Cys -20 -15	
CTC TAT GAT GCC TCC AAC AGC GAG TGG CAG CTC Leu Tyr Asp Ala Ser Asn Ser Glu Trp Gln Leu -5	
TTT CTT CGA GAA TCC TTG GAA CAG AAA CTG GAC Phe Leu Arg Glu Ser Leu Glu Gln Lys Leu Glu 10 15 20	Lys Arg Arg Glu Glu
GAA GTG AGC CAC GCT GGG	364

30

(vi) ORIGINAL SOURCE:

(ix) FEATURE:

(A) ORGANISM: Homo Sapiens(F) TISSUE TYPE: Brain

(A) NAME/KEY: sig\_peptide (B) LOCATION: 103..141

(D) OTHER INFORMATION: score 3.6

(C) IDENTIFICATION METHOD: Von Heijne matrix

(2) INFORMATION FOR SEQ ID NO: 163:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 185 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: CDNA	
<pre>(vi) ORIGINAL SOURCE:     (A) ORGANISM: Homo Sapiens     (F) TISSUE TYPE: Brain</pre>	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 129173     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 3.6</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 163:	
AGTAGACCGC GCACTGGAAG GCGTGCGCGC AGGTGTGCGT GACCATGTGC TTGAAACGGC	60
AGTAGCGCAS RNGNAAGGAT CGCCATCACA CGGCGCACTG GTGCGGCTTC TCCCCCGAGT	120
GGACGAAC ATG TGC TTG GTG TCG TTT TTC CTT GAG CTG AAC GTC TTG CAA  Met Cys Leu Val Ser Phe Phe Leu Glu Leu Asn Val Leu Gln -15 -10 -5	170
CAG TGG CCG GCA GGG Gln Trp Pro Ala Gly 1	185
(2) INFORMATION FOR SEQ ID NO: 164:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 234 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: CDNA	

## seq MRSLACLTPCGHA/GS

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 164:

ATAC	CTCT	TTC (	CAGTT	rggg <i>i</i>	AA AA	AAATO	GACT	TAT 1	AAAT	STCC	CAT	GTCC	AGG (	CTGA	CCTGG	A 6	60
TGAT	GAC	AGT :	rggti	rgat(	GA GI	raat?	TTGA	A AC	ATGAC	GCAG		ATG A		Ser :		11	14
			ACT Thr													16	52
			TAC Tyr													21	10
			TCA Ser													23	34

# (2) INFORMATION FOR SEQ ID NO: 165:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 315 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 70..108
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.5

seq MHLLSNWANPASS/RR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 165:

AAGTGGCCAT (	GGCGGATACA	GCGACTACAG	CATCGGCGGC	GGCGGCTAGT	GCCGCTAGCG	60
--------------	------------	------------	------------	------------	------------	----

- CCTCGAGCG ATG CAC CTC CTT TCC AAC TGG GCA AAC CCC GCT TCC AGC AGA 111

  Met His Leu Leu Ser Asn Trp Ala Asn Pro Ala Ser Ser Arg

  -10 -5 1
- CGT CCT TCT ATG GCC GCT TCA GGC ACT TCT TGG ATA TCA TCG ACC CTC

  Arg Pro Ser Met Ala Ala Ser Gly Thr Ser Trp Ile Ser Ser Thr Leu

  5

  159
- GCA CAC TCT TTG TCA CTG AGA GAC GTC TCA GAG AGG CTG TGC AGC TGC Ala His Ser Leu Ser Leu Arg Asp Val Ser Glu Arg Leu Cys Ser Cys

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127										
20 25	30									
TGG AGG ACT ATA AGC ATG GGA CCC TGC GCC CGG Trp Arg Thr Ile Ser Met Gly Pro Cys Ala Arg 35 40										
AGC TCT GGA GTG CAC AGA AAA TCA AGC AGG CTA Ser Ser Gly Val His Arg Lys Ser Ser Arg Leu 50 55 60										
CCA ATG AGA AGA Pro Met Arg Arg	315									
(2) INFORMATION FOR SEQ ID NO: 166:										
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 415 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR  (ii) MOLECULE TYPE: CDNA  (vi) ORIGINAL SOURCE:										
<ul><li>(A) ORGANISM: Homo Sapiens</li><li>(F) TISSUE TYPE: Brain</li></ul>										
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 62133     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 3.5</pre>										
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:	166:									
AAAGGAGCCA ACYKCACAGT ACATTTCTCT TTGTCATGAA	TTGCATACTT TGTTCCAAGT 60									
C ATG TGG TCT GGA AAG TGG GCG TTG GTC TCA Commet Trp Ser Gly Lys Trp Ala Leu Val Ser Properties Comment of the										
TCA GTG TGG AGA CTC ATT CCT GCC TTT CGT GGT Ser Val Trp Arg Leu Ile Pro Ala Phe Arg Gly -5										
GCT CAG GAA TTT CTT TGT GAA CTT TTA GAT AAA Ala Gln Glu Phe Leu Cys Glu Leu Leu Asp Lys										

GAG ACA ACT GGT ACC AGT TTA CCA GCT CTT ATC CCC ACT TCT CAA AGG

Glu Thr Thr Gly Thr Ser Leu Pro Ala Leu Ile Pro Thr Ser Gln Arg

AAA CTC ATC AAA CAA GTT CTG AAT GTT GTA AAT AAC ATT TTT CAT GGA

Lys Leu Ile Lys Gln Val Leu Asn Val Val Asn Asn Ile Phe His Gly 50

30

45

253

301

W	(U 9)	//0002	12		128									. 1/115/0/		
CAA ( Gln )																349
ACC I																397
CAA :							•									415
(2)					_											
	(i	) SE	(A) (B) (C)	LENG TYPE STRA	CHARA STH: C: NU ANDED OLOGY	252 ICLEI NESS	base C AC S: DC	pai CID OUBLE								
	(i	i) M	OLEC	ULE	TYPE	:: C[	ANC									
	(∨	·i) C	(A)	ORGA	SOUF NISM SUE T	l: Ho		•	ens							
	(i	×) F	(B) (C)	NAME LOCA IDEN	:/KEY TION TIFI CR IN	: 13 CATI	301 [ON 1	189 1ETHO	D: \ scoi	/on l re 3. KFCI	. 5					
	( x	i) S	EQUE	ENCE	DESC	RIPT	rion:	: SE(	Q ID	NO:	167	:				
AAGA	CGC	scc o	GTT	CTG	CG AC	CGCA	GTTA	G CG	CAGT	CTGC	TTT	GGTG	AAT I	ACAC	GATTT	G 60
GTGC	AGCC	CGG (	GTT	rggt	AC CO	GAGC	GGAG	A GG	AGAT	GCAC	ACG	GCAC	TCG I	AGTG'	TGAGG	A 120
AAAA	TAGA	Me					et H					ys L			GT TT ys Le	
CTG . Leu																219
CAT His										Gly						252
(2)	TNE	מאפר	TION	FOR	SEO	TD	NO ·	168.								

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 436 base pairs
    (B) TYPE: NUCLEIC ACID

	•=-	
(C)	STRANDEDNESS: DOUBLE	
(D)	TOPOLOGY: LINEAR	

- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 290..361
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.5

seq ALSLFYTADTSHG/SE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 168:

ATATTTCTTG	TCAACAGTAT	TGAAATGTAA T	ATGTATGTG TT	CATGTATG AGMA	ATTTTT 60
ACTCCACACA	GGTGTTTCAG	TAGAGTGGGG C	AGGAAAAGA GA	ATCTCTTCG ATTT	CTTTCA 120
GGCCTGAGGC	TTTTGTGAAA	TGCGTCASCC C	CTGTGACAG TA	AGGTTTTGA TGCT	AGTGAT 180
CTTCAGATCT	TTCTCTCTGG	AAATGTGCAG A	GAGTGTCAG TI	TTCCCAAGT TCTG	AGGTAA 240
CTCTCAGCCC	AGATGTGAAA	TGGGAGCCTA C	CAGCTGGTA TA	AGAAGGGA ATG G Met G	GT AGG 298 ly Arg
			a Leu Ser Le	TT TTC TAT ACT eu Phe Tyr Thr 10	
				GT CTA TTT GGA er Leu Phe Gly 10	Arg
		GR AGT AAC CC ly Ser Asn Pr 2			436

- (2) INFORMATION FOR SEQ ID NO: 169:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 343 base pairs

    - (B) TYPE: NUCLEIC ACID
      (C) STRANDEDNESS: DOUBLE
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: CDNA
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: other
    - (B) LOCATION: 104..336

O 99/06552

(C) IDENTIFICATION METHOD: blastn
(D) OTHER INFORMATION: identity 96
region 1..233

id H07998

## (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 110..336
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 96

region 1..227 id W37530 est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 110..336
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 96

region 1..227 id R79812 est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 110..336
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 96

region 1..227 id N24900 est

### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 110..336
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 96

region 1..227 id R34849 est

## (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 65..112
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 12.5

seq FVVLLALVAGVLG/NE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 169:

ATGTCGCCCG TGTCCCGCCG GCCCGTTCCG TGTCGCCCCG CAGTGYTGCG GCCGCCGCKK 60

GGG AAC GAG TTT AGT ATA TTA AAA TCA CCA GGG TCT GTT GTT TTC CGA
Gly Asn Glu Phe Ser Ile Leu Lys Ser Pro Gly Ser Val Val Phe Arg
1 5 10

AAT GGA AAT TGG CCT ATA CCA GGA GAG CGG ATC CCA GAC GTG GCT GCA 205 Asn Gly Asn Trp Pro Ile Pro Gly Glu Arg Ile Pro Asp Val Ala Ala 20 TTG TCC ATG GGC TTC TCT GTG AAA GAA GAC CTT TCT TGG CCA GGA CTC 253 Leu Ser Met Gly Phe Ser Val Lys Glu Asp Leu Ser Trp Pro Gly Leu GCA GTG GGT AAC CTG TTT CAT CGT CCT CGG GCT AGC GTC ATG GTG ATG 301 Ala Val Gly Asn Leu Phe His Arg Pro Arg Ala Ser Val Met Val Met 50 55 GTG AAG GGA GTT AAC AAC TMC CCT CTA CCC CCA NGN TGG NGG 343 Val Lys Gly Val Asn Asn Xaa Pro Leu Pro Pro Xaa Trp Xaa 70

#### (2) INFORMATION FOR SEQ ID NO: 170:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 234 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 111..209
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 90

region 1..99 id N50844

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 186..232
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 93

region 75..121

id N50844

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 111..209
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 90

region 1..99

id N29905

est

(ix) FEATURE:

WO 99/06552 PCT/IB98/01236

(A) NAME/KEY: other (B) LOCATION: 186232 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 93 region 75121 id N29905 est	
(ix) FEATURE:  (A) NAME/KEY: other  (B) LOCATION: 186232  (C) IDENTIFICATION METHOD: blastn  (D) OTHER INFORMATION: identity 93  region 75121  id N62597  est	
(ix) FEATURE:  (A) NAME/KEY: other  (B) LOCATION: 186232  (C) IDENTIFICATION METHOD: blastn  (D) OTHER INFORMATION: identity 93  region 76122  id R80247  est	
(ix) FEATURE:  (A) NAME/KEY: other  (B) LOCATION: 186232  (C) IDENTIFICATION METHOD: blastn  (D) OTHER INFORMATION: identity 93  region 76122  id H03409  est	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 4087     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 10.1</pre>	
AAGAGGTGCG GGATTGGGCG GGCTGCCACG GCATGGAGA ATG GCT CCG CTT CTG Met Ala Pro Leu Leu -15	54
TTG CAG CTG GCG GTG CTC GGC GCG GCG CTG GCG GC	102
CTG ATT TCC ATC GTT GCA TTT ACA ACT GCT ACA AAA ATG CCA GCA CTC Leu Ile Ser Ile Val Ala Phe Thr Thr Ala Thr Lys Met Pro Ala Leu 10 15 20	159
CAT CGA CAT GAA GAA GAG AAA TTC TTC TTA AAT GCC AAA GGC CAG AAA His Arg His Glu Glu Lys Phe Phe Leu Asn Ala Lys Gly Gln Lys 25 30 35	193

133 GAA ACT TTA CCC AGC ATA TGG GAC TCA CCT ACC AGG 234 Glu Thr Leu Pro Ser Ile Trp Asp Ser Pro Thr Arg 40 45 (2) INFORMATION FOR SEQ ID NO: 171: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 386 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR (ii) MOLECULE TYPE: CDNA (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 52..228 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 94 region 1..177 id AA074050 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 266..387 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 99 region 218..339 id AA074050 est (ix) FEATURE: (A) NAME/KEY: sig\_peptide (B) LOCATION: 135..284 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 9.8 seq LLRLLQLVSTCVA/FS (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 171: AGCGGCCGCA GCCAGCCAGG CCGCGCMMGG GACGACTGCA GAGCGCGGTG CTCTTACAGC 60

AGCGGCCGCA GCCAGCCAGG CCGCGCMMGG GACGACTGCA GAGCGCGGTG CTCTTACAGC

CTGTTCCAAG TGTGGCTTAA TCCGTCTCCA CCACCAGATC TTTCTCCGTG GATTCCTCTG

CTAAGACCGC TGCC ATG CCA GTG ACG GTA ACC CGC ACC ACC ATC ACA ACC

Met Pro Val Thr Val Thr Arg Thr Thr Ile Thr Thr

-50

ACC ACG ACG TCA TCT TCG GGC CTG GGG TCC CCC ATG ATC GTG GGG TCC

Thr Thr Thr Ser Ser Ser Gly Leu Gly Ser Pro Met Ile Val Gly Ser

-35

CCT CGG GCC CTG ACA CAG CCC CTG GGT CTC CTT CGC CTG CAG CTG

266

Pro Arg Ala Leu Thr Gln Pro Leu Gly Leu Leu Arg Leu Leu Gln Leu

GTG TCT ACC TGC GTG GCC TTC TCG CTG GTG GCT AGC GTG GGC GCC TGG 314 Val Ser Thr Cys Wal Ala Phe Ser Leu Val Ala Ser Val Gly Ala Trp

ACG GGG TCC ATG GGC AAC TGG TCC ATG TTC ACC TGG TGC TTC TGC TTC Thr Gly Ser Met Gly Asn Trp Ser Met Phe Thr Trp Cys Phe Cys Phe 15 20

TCN GTG ACC CTG ATC ATC CTC ATC Ser Val Thr Leu Ile Ile Leu Ile

386

### (2) INFORMATION FOR SEQ ID NO: 172:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 326 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

#### (ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

### (ix) FEATURE:

(A) NAME/KEY: other

(3) LOCATION: 147..290

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99 region 57..200 id W40499

est

## (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 90..151

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98 region 1..62 id W40499

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 100..319

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 46..265

id R88049

est

### (ix) FEATURE:

(A) WAME/KEY: other

(B) ECCATION: 100..319

(C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 99 region 56..275 id T08712 (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 100..319 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 99 region 32..251 id H38484 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 147..319 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 98 region 65..237 id T65344 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 102..151 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 100 region 21..70 id T65344 est (ix) FEATURE: (A) NAME/KEY: sig\_peptide (3) LOCATION: 111..164 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 9.2 seq VFLCSLLAPMVLA/SA (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 172: AGACTCTTGG GGACTGGGCT GAGGACGGGG TGGTACTGCT CCTGGCAGGG CCAGAGGTGG ATGGGGCTTG AAAAGGGGGT TCAAGGCAGC AGMTCTATGG TTCAGACGCC ATG GAG 116 Met Glu TTG GTG CTG GTC TTC CTC TGC AGC CTG GCC CCC ATG GTC CTG GCC 164 Leu Val Leu Val Phe Leu Cys Ser Leu Leu Ala Pro Met Val Leu Ala -15 AGT GCA GCT GAA AAG GAG AAG GAA ATG GAC CCT TTT CAT TAT GAT TAC 212 Ser Ala Ala Glu Lys Glu Lys Glu Met Asp Pro Phe His Tyr Asp Tyr 10 CAG ACC CTG AGG ATT GGG GGA CTG GTG TTC GCT GTG GTC CTC TTC TCG 260 Gln Thr Leu Arg Ile Gly Gly Leu Val Phe Ala Val Val Leu Phe Ser 20 GTT GGG ATC STS STT ATC CTA AGT CGC AGG TGC AAG TGC AGT TTC AAT

Val Gly Ile Leu Leu Ile Leu Ser Arg Arg Cys Lys Cys Ser Phe Asn 35 40 45

CAG AAG CCC CGC AAC AGA Gln Lys Pro Arg Asn Arg 50 326

- (2) INFORMATION FOR SEQ ID NO: 173:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 376 base pairs
    - (B) TYPE: NUCLEIC ACID
    - (C) STRANDEDNESS: DOUBLE
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: CDNA
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: other
    - (B) LOCATION: 74..344
    - (C) IDENTIFICATION METHOD: blastn
    - (D) OTHER INFORMATION: identity 96

region 73..343 id H95186

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 25..86
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 93 region 25..86

id H95186

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 138..377
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 100

region 1..240

id N40665

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 203..308
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 95

region 230..335

id W25197

est

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide

WO 331003.	J4			101,127								
	(C) IDE	NTIFICAT	: 167304 CATION METHOD: Von Heijne matrix FORMATION: score 7.9 seq LLGLLSAEQLAEA/SV									
(xi)	SEQUENCE	DESCRIPT	TION: SEC	Q ID NO:	173:							
AACGGGCGTC	GGGAAGAC	GC TGCTG	GTGAA AC	GCTGCAG	GAGGTGAG	GCT CCCG	GGATGG	60				
GAAAGGCGAC CTGGGGGAGC CGCCCCCGAC ACGGCCCACG GTGGGCACCA ATCTTACTGA 1												
CATCGTGGCA	CAGAGAAA	GA TCACC	ATCCG GG	AGCTTGGG		ATG GGC ( Met Gly :		.75				
ATC TGG TCC Ile Trp Ser								223				
GAC GCC TCT Asp Ala Sec -25	Asp Pro							271				
GGT CTC CTT Gly Leu Leu -10								319				
TTC AAT AAP Phe Asn Lys		Leu Pro					Lys	367				
TCA TTA ATO Ser Leu Ile	-							376				
(2) INFORM	ATION FOR	SEQ ID	NO: 174:									
(i) \$	(B) TYP (C) STR	CHARACTE GTH: 277 E: NUCLE ANDEDNES: OLOGY: L	base pa IC ACID S: DOUBL									
(ii)	MOLECULE	TYPE: C	DNA									
(vi)		SOURCE: ANISM: H SUE TYPE		ens								
(ix)	(B) LOC	E/KEY: o ATION: 5 NTIFICAT ER INFOR	9278 ION METH	OD: blas identit region id R789	y 97 28247							

id R78970 est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 59..210

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 92

region 29..180 id R64509

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 196:.278

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 167..249 id R64509

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 59..210

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 92

region 44..195

id H85714

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 196..278

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 182..264

id H85714

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 59..278

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 36..255

id H52756

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 59..278

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96 region 5..224

id H49758

(im) FEATURE:

(A) NAME/KEY: sig\_peptide.

(B) LOCATION: 107..247

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 7.9

seq LLLPRVLLTMASG/SP

(xil REQUENCE DESCRIPTION: SEQ ID NO: 174:

139

ATC	ACGTO	GC 1	RGCC	ACCC	AG GI	KAMGA	AAGAI	R NAI	NVTC1	rtcc	TGG	GTTV	/HT '	rctc	CGANRT	60
GAC	STSYS	SGC (	CTTT	GAGAT	rc a <i>l</i>	ACTCT	rcct(	G TAC	CCAGO	CGTA	GGC			Ser (		115
			CCC Pro													163
			TGG Trp -25													211
			CGA Arg													259
			CCG Pro													277
(2)	TNEC	ימשפר	TTON	FOR	SEO.	TD N	10. 1	175.								

#### (2) INFORMATION FOR SEQ ID NO: 175:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 388 base pairs (B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 180..390

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98 region 134..344

id H08480

est

(ix) FEATURE:

(A) NAME/KEY: other (3) LOCATION: 115..185

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 1..71 id H08480

est

(im; FEATURE:

(A) NAME/KEY: sig\_peptide (B) LOCATION: 113..232

. (C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 7.9

seq SLLLLFGGQFASS/QE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 175:

AGCAGAGCTT CCGCTTCCGG CCCTTCAGGC TCTGTCTCTG TGGAGACTGG GCTTTGGGAG 60 GKAGAAAGAG GGACCTAGCG CGGGCCGCGC AGGCGCACGG TGGGCAGCTG CA ATG GCG Met Ala CTG TCG TGT ACC CTT AAC AGG TAT CTG CTC CTC ATG GCG CAG GAG CAT Leu Ser Cys Thr Leu Asn Arg Tyr Leu Leu Leu Met Ala Gln Glu His -35 -30 CTG GAG TTC CGC CTG CCG GAA ATA RRG TCT TTG CTT TTG CTT TTT GGA Leu Glu Phe Arg Leu Pro Glu Ile Xaa Ser Leu Leu Leu Phe Gly -15 GGT CAG TTT GCC AGC AGT CAA GAA ACT TAT GGA AAG TCA CCA TTT TGG 262 Gly Gln Phe Ala Ser Ser Gln Glu Thr Tyr Gly Lys Ser Pro Phe Trp ATT CTT AGC ATT CCC TCT GAA GAT ATT GCA AGA AAT TTG ATG AAA CGG Ile Leu Ser Ile Pro Ser Glu Asp Ile Ala Arg Asn Leu Met Lys Arg 15 ACA GTG TGT GCC AAG TCT ATA TTT GAA CTA TGG GGT CAT GGA CAA TCT 358 Thr Val Cys Ala Lys Ser Ile Phe Glu Leu Trp Gly His Gly Gln Ser 30

CCT GAG GAG CTG TAC AGT TCT CTT AAA AAC Pro Glu Glu Leu Tyr Ser Ser Leu Lys Asn 45 388

## (2) INFORMATION FOR SEQ ID NO: 176:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 311 base pairs
  - (3) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (im) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 112..309
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 97 region 69..266 id AA149265

est

## (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 41..86
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 97

region 1..46 id AA149265

est

## (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 110..309
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 97

region 53..252 id W39570

est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 56..86
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 96

region 2..32 id W39570 est

## (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 110..309
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 97

region 55..254

id N41332

est

## (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 53..86
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 97

region 1..34 id N41332

est

## (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 39..197
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 7.1

seq IAVGLGVAALAFA/GR

(x1) SEQUENCE DESCRIPTION: SEQ ID NO: 176:

AACTGUUGUU GEGEETTGAG TETEEGGGEE GEETTGEE ATG GET GEE GGT GTC 56
Met Ala Ala Arg Gly Val

-50

Ile Ala Pro Val Gly Glu Ser Leu Arg Tyr Ala Glu Tyr Leu Gln Pro -40 TCG GCC AAA CGG CCA GAC GCC GAC GTC GAC CAG CAG AGA CTG GTA AGA Ser Ala Lys Arg Pro Asp Ala Asp Val Asp Gln Gln Arg Leu Val Arg AGT TTG ATA GCT GTA GGA CTG GGT GTT GCA GCT CTT GCA TTT GCA GGT 200 Ser Leu Ile Ala Val Gly Leu Gly Val Ala Ala Leu Ala Phe Ala Gly -15 -10 CGC TAC GCA TTT CGG ATC TGG AAA CCT CTA GAA CAA GTT ATC ACA GAA Arg Tyr Ala Phe Arg Ile Trp Lys Pro Leu Glu Gln Val Ile Thr Glu 10 ACT GCA AAG AAG ATT TCA ACT CCT AGC TTT TCA TCC TAC TAT AAA GGA 296 Thr Ala Lys Lys Ile Ser Thr Pro Ser Phe Ser Ser Tyr Tyr Lys Gly 25 GGA TTT GAA CGG AGG 311 Gly Phe Glu Arg Arg 35

## (2) INFORMATION FOR SEQ ID NO: 177:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 384 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 43..87
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 100 region 8..52

id W32101

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 89..129
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 100 region 53..93

id W32101

et.

- (im) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 292..375
  - (C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 6.6 seq VLGXLFLGGLCRG/WD

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 177:

AAGAGCCGCG TTYAGTCTAT CGCTGCGGTT GCGAGCGCTG TAGGGAGCCT GTGCTGTGCC 60 GCGCAGTTAG GCAGCAGCAG CCGCGGAGCA GTAGCCGCCG TGGGAGGGAG CCATGAAGCA 120 TTACGAGGTA AGAAGCGAGA AACAGGGGCC GTGTGGCCAC TGCTGACCCA TTCTTTTCC TTCTTTGCGG GACCACGGGA CCCCACTTTC TGGTCCTGTG CCCCGAAGGA AGAKCCAGAC GGCGCAGGCG CAGTGGGCAA GCGTTGCGCC CCGGGCCACT CGTAAATTCC A ATG CGC 297 Met Arg ATG TGC GCA GGA AGT ATT TAT AAA TCT GCA ACC CAG GCT GTT TTG GGG 345 Met Cys Ala Gly Ser Ile Tyr Lys Ser Ala Thr Gln Ala Val Leu Gly GWA CTT TTT CTT GGG GGT CTC TGC AGG GGC TGG GAC GCT 384 Xaa Leu Phe Leu Gly Gly Leu Cys Arg Gly Trp Asp Ala -5

#### (2) INFORMATION FOR SEO ID NO: 178:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 425 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 73..317

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 1..245 id HUM506F10B

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 314..376

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98 region 243..305

id HUM506F10B

est

- (ix) FEATURE:
  - (A) NAME/KEY: other

(B) LOCATION: 63..193

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 1..131 id AA056148 est

#### (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 314..401

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 254..341 id AA056148

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 277..317

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 216..256 id AA056148

est

#### (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 397..426

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 338..367 id AA056148

est

### (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 88..189

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 1..102 id HSC1FF051

est

# (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 314..401

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 230..317 id HSC1FF051

est

# (ix) FEATURE:

(A) NAME/KEY: other(3) LOCATION: 187..271

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 101..185 id HSC1FF051

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 269..317

(C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 93

region 184..232

id HSC1FF051

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 397..426

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 314..343 id HSC1FF051

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 87..200

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 92

region 1..114 id HSC16E081 est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 314..401

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 231..318 id HSC16E081

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 199..275

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 114..190 id HSC16E081

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 269..317

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 91

region 185..233 id HSC16E081

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 397..426

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 315..344 id HSC16E081

identity 99 region 24..125 id AA157365

identity 90 region 123..203 id AA157365

•		146
		est
(ix)	FEATURE: (A) NAME/KEY: other (B) LOCATION: 85186 (C) IDENTIFICATION ME (D) OTHER INFORMATION	ETHOD: blastn
(ix)	FEATURE: (A) NAME/KEY: other (B) LOCATION: 18326 (C) IDENTIFICATION ME (D) OTHER INFORMATION	ETHOD: blastn
(ix)	FEATURE: (A) NAME/KEY: other (B) LOCATION: 33740 (C) IDENTIFICATION ME	

- HOD: blastn
- (D) OTHER INFORMATION: identity 95

region 278..342 id AA157365

- (ix) FEATURE:
  - (A) NAME/KEY: other (B) LOCATION: 273..326
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 90 region 213..266 id AA157365

est

(ix) FEATURE:

-60

- (A) NAME/KEY: sig\_peptide (B) LOCATION: 186..419
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 6.3

seq TLIMLLSWQLSVS/SV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 178:

AATGCGCGCT CGCGNTCCCG CCCTCTAGCT GCGCTCGGCT GAGTCAGTCA GTCTGTCGGA 60 STCTGTCCTC GGAGCAGGCG GAGTAAAGGG ACTTGAGCGA GCCAGTTGCC GGATTATTCT ATTTCCCCTC CCTCTCCCC GCCCGTATC TCTTTTCACC CTTCTCCCAC CCTCGCTCGC 180 STASC ATG GCG GAG CGT CGG CGG CCA CTC AGT CCC ATT CCA TCT NNT CGT Met Ala Glu Arg Arg Pro Leu Ser Pro Ile Pro Ser Xaa Arg -75 COT CCT TCG GAG CCG AGC CGT CCG CGC CCG GCG GCG GCG GGA SCC AGG 273 Ang Pro Ser Glu Pro Ser Arg Pro Arg Pro Ala Ala Ala Gly Xaa Arg

-55

AGC CTG CCC CGC CCT GGG GAC GAA GAG CTG CAG CTC CCC TGT GCG GTG Ser Leu Pro Arg Pro Gly Asp Glu Glu Leu Gln Leu Pro Cys Ala Val -45

CAC GAT CTG ATT TTC TGG AGA GAT GTG AAG AAG ACT GGG TTT GTC TTT Arg Asp Val Lys Lys Thr Gly Phe Val Phe -20

GGC ACC ACG CTG ATC ATC ATG CTG CTT TCC TGG CAG CTT TCA GTG TCA TCA GGG TTT Thr Leu Ile Met Leu Leu Ser Trp Gln Leu Ser Val Ser Ser -15

GTG Val

### (2) INFORMATION FOR SEQ ID NO: 179:

#### (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 403 base pairs
- (B) TYPE: NUCLEIC ACID
- (C) STRANDEDNESS: DOUBLE
- (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 105..351
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 98

region 49..295

id R47336

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 58..107
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 98

region 1..50

id R47336

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 352..381
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 100

region 295..324

id R47336

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide

(B) LOCATION: 5..331

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 6

seq LQLLLGMTASAVA/AL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 179:

		AGA GTG TCG GTG CCG Arg Val Ser Val Pro -100	
	Ala Val Cys A	GCT GCC GGA ATC CTG Ala Ala Gly Ile Leu -85	
	Glu Arg Glu I	AAG GAG CGG GAC CCC Lys Glu Arg Asp Pro -70	
		GTG AAG TGC TCC GCC /al Lys Cys Ser Ala -50	
		CTT TTG GGT TCC ATT Leu Leu Gly Ser Ile -35	
		AGT GTC TTT GGA CTT Ser Val Phe Gly Leu -20	
	Leu Gly Met	ACA GCA AGC GCT GTG Thr Ala Ser Ala Val -5	
		TCG GTC GTG GGG TCC Ser Val Val Gly Ser 15	
CCT ACA TTC TGT ACT Pro Thr Phe Cys Thr 20			403

# (2) INFORMATION FOR SEQ ID NO: 180:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 367 base pairs
  - (B) TYPE: NUCLEIC-ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 112..260

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 92

region 121..269 id W31320

est

(ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 47..118

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 57..128

id W31320

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 273..333

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 282..342

id W31320

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 107..260

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 2..155

id T27259

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 273..369

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 168..264

id T27259

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 145..260

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 90

region 108..223

id AA157646

est

(ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 59..118

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 90

region 25..84

id AA157646

	(i	x) F	(B) (C)	RE: NAME LOCA IDEN OTHE	TION TIFI	: 27 CATI	33 ON M	ETHO	iden	tity on 2	94 45	279			
	·	·	(B) (C) (D)	NAME LOCA I DEN OTHE	TION TIFI R IN	: 50 CATI FORM	15 ON M	1 IETHC ON:	D: V scor seq	e 5. LGAA	9 ALAL	LLAN			
	( x	:i) S	EQUE	NCE	DESC	RIPT	'ION:	SEC	DI	NO:	180:				
\AT <i>F</i>	ATACI	TC I	TTTGT	CAAG	SA GA	AGCA	GAGO	G TGT	rggac	GCT	GTG1	TATGA		T TTC	58
				AGT Ser											106
				GCT Ala											154
				AAG Lys											202
				ACA Thr											250
				AAA Lys											298
				TGT Cys											346
				CAG Gln 70											367
(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	181:							

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 257 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 138..257

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 83..202

id W31692

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 55..131

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 1..77 id W31692

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 136..257

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 78..199 id H50194

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 57..131

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 1..75 id H50194

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 57..257

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 1..201

id H46855

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 138..257

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 81..200

id H49687 est

(ix) FEATURE:

152 . (A) NAME/KEY: other (B) LOCATION: 57..132 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 100 region 1..76 id H49687 (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 138..257 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 98 region 80..199 id T54405 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 58..124 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 100 region 2..68 id T54405 est (ix) FEATURE: (A) NAME/KEY: sig\_peptide (B) LOCATION: 90..200

- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 4.9

seq MLIMLGIFFNVHS/AV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 181:

ATCTCTGCCC CCCTGCGAGG GCATCCTGGG CTTTCTCCCA CCGCTTTCCG AGCCCGCTTG 60 CACCTCGGCG ATCCCCGACT CCCTTCTTT ATG GCG TCG CTC CTG TGC TGT GGG 113 Met Ala Ser Leu Leu Cys Cys Gly -35 CCG AAG CTG GCC GCC TGC GGC ATC GTC CTC AGC GCC TGG GGA GTG ATC Pro Lys Leu Ala Ala Cys Gly Ile Val Leu Ser Ala Trp Gly Val Ile -25 ATG TTG ATA ATG CTC GGA ATA TTT TTC AAT GTC CAT TCC GCT GTG TTG 209 Met Leu Ile Met Leu Gly Ile Phe Phe Asn Val His Ser Ala Val Leu -10 ATT GAG GAC GTT CCC TTC ACG GAG AAA GAT TTT GAG AAC GGC CCC CGG 257 Ile Glu Asp Val Pro Phe Thr Glu Lys Asp Phe Glu Asn Gly Pro Arg

# (2) INFORMATION FOR SEQ ID NO: 182:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 400 base pairs
  - (B) TYPE: NUCLEIC ACID

,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,	153	_
	DNESS: DOUBLE	
(D) TOPOLOG	GY: LINEAR	
(ii) MOLECULE TYP	PE: CDNA	

(ix) FEATURE:

(vi) ORIGINAL SOURCE:

- (A) NAME/KEY: other
- (B) LOCATION: 365..401

(A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100 region 1..37 id R50224 est

- (ix) FEATURE:
  - (A) NAME/KEY: sig peptide
  - (B) LOCATION: 305..364
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.9

seq XSLFLHAVSSSFT/QL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 182:

ATGACCACGG GTTTAACCTT CTTATCCCAG AGACACCCAA TTCTAGAGCT TTATGGAGCC 60 GTACTTCCCC CTGAATCCTA GCTCTAGGAC ATAGATCATG ACTCTCAGCC CTTTTACCCA 120 GGATGGAGCT GGGGCCTGTA TAGCCATATT ATTGTTCTAA GTAAGTTCTA GCCCCACCCT 180 CCCGCCTTCT TGAGTGATAC CTATTACGGA TGAGTTCTGG AAAAGACCCA GCTATGATTC 240 ATAAAAACAC TTCTGGATGA ATCAAGAACC ATTTCTTGTT TKTCCTAGAT AATTCTCTAA AAAT ATG ATT CTT CCA TAT AGA ATG CKA AGC TTA TTT TTA CAT GCA GTT Met Ile Leu Pro Tyr Arg Met Xaa Ser Leu Phe Leu His Ala Val -20 -15 TCT AGC TCC TTC ACC CAG CTG AGG TCG TGC CAG GGA GAC AGA GTC TGG 397 Ser Ser Ser Phe Thr Gln Leu Arg Ser Cys Gln Gly Asp Arg Val Trp -5 1 400 AGA Arg

- (2) INFORMATION FOR SEQ ID NO: 183:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 256 base pairs
    - (3) TYPE: NUCLEIC ACID
    - (C) STRANDEDNESS: DOUBLE
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: CDNA

	(v.	i) (		ORGA	SOUR NISM UE T	: Ho			ns						
	(i:	×) I	(B) (C)	NAME LOCA I DEN	/KEY TION TIFI R IN	: 86 CATI	18 ON M	ETHO N:	iden regi	tity	93 10	5			
	·		(B) (C) (D)	NAME LOCA IDEN OTHE	/KEY TION TIFI R IN	: 23 CATI	ON M	1 IETHO N:	D: V scor seq	e 4. LYTV	8 'RALA	GRAW	trix NA/AV		
	(x	i) :	SEQUE	NCE	DESC	RIPT	CION:	SEC	) ID	NO:	183:				
AGA!	AGTGC	GT :	SYCGO	CGGG	SA TO					ı Val				TCG Ser	52
	CTG Leu														100
	TCC Ser		Val												148
	TGG Trp -20														196
	CGG Arg														244
	CCA Pro														256
(2)	INFO	RMA	TION	FOR	SEQ	ID	NO:	184:							
	(i	.) S	(B) (C)	LENG TYPE STR	CHARA GTH: E: NO ANDEI OLOG	352 UCLE DNES	base IC AG S: DG	e pa: CID DUBLI							
	( i	i)	MOLE	CULE	TYP	E: C	DNA								
	(1	7 <u>i</u> )	ORIG	INAL	SOU	RCE:									

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

### (ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 183..348

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 78..243 id W52941

est

# (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 286..348

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 1..63 id H55390

### (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 77..199

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.7

seq LFSCFCFLSHKFG/KK

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 184:

# AAAAAATATC TCCCGGCGTG CGCTGCTTGT GTTATGTTCG GGTTTTAAGT CGTGTCAGCG 60

TTTACATTTT CTTAAT ATG AAA AAT GCC TGC ATT GTT CTG CCG CCA ACT CCC 112

Met Lys Asn Ala Cys Ile Val Leu Pro Pro Thr Pro

-40 -35 -30

CCT CCC TCC CTG CAA CCC TCG GCC TCT CTG CTG GCG CCT AAT CGT TTT

Pro Pro Ser Leu Gln Pro Ser Ala Ser Leu Leu Ala Pro Asn Arg Phe

-25

-15

TTA TTC TCT TGC TTC TGC TTT CTT AGT CAC AAG TTT GGG AAG AAA GTC

Leu Phe Ser Cys Phe Cys Phe Leu Ser His Lys Phe Gly Lys Lys Val

-10

-5

ATC TAT TTC AAC TAC CTG AGT GAG CTC CAC GAA CAC CTT AAA TAC GAC

1le Tyr Phe Asn Tyr Leu Ser Glu Leu His Glu His Leu Lys Tyr Asp

10

15

CAG CTG GTC ATC CCT CCC GAA GTT TTG CGG TAC GAT GAG AAG CTC CAG
Gln Leu Val Ile Pro Pro Glu Val Leu Arg Tyr Asp Glu Lys Leu Gln
20 25 30 35

AGC CTG CAC GAG GGC CGG ACG CCG MCT CCC ACC AAG ACA CCA CCA GGG 352 Ser Leu His Glu Gly Arg Thr Pro Xaa Pro Thr Lys Thr Pro Pro Gly

### (2) INFORMATION FOR SEQ ID NO: 185:

(i) SEQUENCE CHARACTERISTICS:

. (A) LENGTH: 274 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

# (ii) MOLECULE TYPE: CDNA

# (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 99..260

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 126..287

id T53519

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 40..108

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 91

region 1..69 id T53519

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 113..269

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 131..287

id W87344

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 147..269

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 138..260

id N56542

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 113..149

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 91

region 105..141

id N56542

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 75..105

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

WO 99/06552 PCT/IB98/01236

region 1..31 id N56542

est

#### (ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 113..218

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 117..222 id AA053475

est

### (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 218..269

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 223..274 id AA053475

est

### (ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 113..269

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 90..246 id W05444

est

# (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 110..193

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.6

seq PLQWSLLVAVVAG/SV

### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 185:

ACTTCCGCCT GCGCCTGCGC AGCVCAGCTC CSHGAGCCCT GCCAACCATG GTGAACTTGG GTCTGTCCCG GGTGGACGAC GCCGTGGCTG CCAAGCACCC GGCACCGGC ATG GCC TTT Met Ala Phe GGC TTG CAG ATG TTC ATT CAG AGG AAG TTT CCA TAC CCT TTG CAG TGG 166 Gly Leu Gln Met Phe Ile Gln Arg Lys Phe Pro Tyr Pro Leu Gln Trp -15-25 -20 AGC CTC CTA GTG GCC GTG GTT GCA GGC TCT GTG GTC AGC TAC GGG GTG 214 Ser Leu Leu Val Ala Val Val Ala Gly Ser Val Val Ser Tyr Gly Val ACG AGA GTR RAG TCG GAG AAA TGC AAC AAC CTC TGG CTC TTC CTG GAG Thr Arg Val Xaa Ser Glu Lys Cys Asn Asn Leu Trp Leu Phe Leu Glu 10 15 ACC GGA CTT GGG 274

Thr Gly Leu Gly 25

WO 99/06552

(2) INFORMATION FOR SEQ ID NO: 186:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 316 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR	
(ii) MOLECULE TYPE: CDNA	
<pre>(vi) ORIGINAL SOURCE:    (A) ORGANISM: Homo Sapiens    (F) TISSUE TYPE: Brain</pre>	
(ix) FEATURE:  (A) NAME/KEY: other  (B) LOCATION: 45315  (C) IDENTIFICATION METHOD: blastn  (D) OTHER INFORMATION: identity 98  region 1271  id HSC1ZD051  est	
<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 110268     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 4.5</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 186:	
ATATGAGACT CTGGCCTCCC TGCAGATCTT CTAAGAACCA CACTAATGCA AGCGTGACAG	60
AGAAACCTCT TTCGAATGAC CTACTACAAC TCTGGCATTG GTTAGTTCC ATG TAT TGT Met Tyr Cys	118
AAG ATT CTG GTG CTA ATG CTC CAT ACA GAA TTG ATC AGG ACT GAT TAC Lys Ile Leu Val Leu Met Leu His Thr Glu Leu Ile Arg Thr Asp Tyr -50 -45 -40 -35	166
TCT TCT GTG GAC CAA TTG CTA TTG AAC TAC CCA GCT GAA GAG GGT TTG Ser Ser Val Asp Gln Leu Leu Leu Asn Tyr Pro Ala Glu Glu Gly Leu -30 -25 -20	214
GGG AGA GAA CGT TCA TTA TTA TGG ACT CCA CTT TTG TCS CCT GGT AGT Gly Arg Glu Arg Ser Leu Leu Trp Thr Pro Leu Leu Ser Pro Gly Ser -15 -10 -5	262
TTA AGG GTG ATA CTA GAA TCC AGA GAA GTT CCT GTC TCC TTG TGG CCC Leu Arg Val Ile Leu Glu Ser Arg Glu Val Pro Val Ser Leu Trp Pro 1 5	310
CAA ACG Gln Thr	31

### (2) INFORMATION FOR SEQ ID NO: 187:

### (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 423 base pairs
- (B) TYPE: NUCLEIC ACID
- (C) STRANDEDNESS: DOUBLE
- (D) TOPOLOGY: LINEAR

### (ii) MOLECULE TYPE: CDNA

### (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo Sapiens
- (F) TISSUE TYPE: Brain

# (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 50..246
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 96

region 1..197 id AA043070

est

# (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 241..373
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 97

region 191..323 id AA043070

est

# (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 371..408
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 100

region 322..359 id AA043070

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 186..357
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 95

region 29..200

id W81202

est

### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 345..423
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 100

region 189..267 id W81202

	(i	.x). E	(B) (C)	NAME LOCA I DEN	TION	: ot : 64 CATI	17 ON M	ETHC	iden regi	olast stity on 8 2485	100 51					
	(i	.x) E	(3) (C)	NAME LOCA IDEN	TION	: ot : 17 CATI	82 ON M	ETHO	iden regi	plast stity on 1 12485	90 98	247				
	<b>(</b> i	. <b>x)</b>	(B) (C)	NAME LOCA IDEN	TION	: si I: 16 CATI IFORM	62 ON M	43 ETHO	D: V	on H e 4. ENSL	5					
	( x	i) S	EQUE	NCE	DESC	RIPT	NOI:	SEC	ÒİD	NO:	187:					
AAC1	CTGC	GC (	CCGGF	AGGAC	CA GA	AGCGG	cccc	GTO	GCCG	GCA	TGGI	TTCT	cc c	STCCI	GCTGC	60
AGC	CGGCG	GG A	AGGC	AGCCF	G TO	CAG	CGCC	CGC	CTAGO	CTTC	GGCG	GCGF	cc c	CAGAC	GGGGA	120
AAGO	CGGAA	AGG P	atg1	CGCC	ST GC	CAAGO	CAGGO	AGC	TGGT	TGTG	GAAC			la Va	G AGC	_
	TCA Ser															225
	CAG Gln -5															273
	GTG Val															321
	CTG Leu															369
	GAT Asp		Leu					Arg								417

423

CCA GAT

Pro Asp 60 (2) INFORMATION FOR SEQ ID NO: 188: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 343 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR (ii) MOLECULE TYPE: CDNA (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 165..302 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 93 region 33..170 id T50032 est (ix) FEATURE: (A) NAME/KEY: other (3) LOCATION: 291..339 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 97 region 160..208 id T50032 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 132..172 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 97 region 1..41 id T50032 est (ix) FEATURE: (A) NAME/KEY: sig\_peptide (B) LOCATION: 71..139 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 4.4 seq QFILLGTTSVVTA/AL (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 188: AAGGTGGAGA GTCGGGGGTC ACCAGGCCTA TCCTTGGCGC CACAGTCGGC CACCGGGGCT 60 CGCCGCCGTC ATB GAG AGC GGA GGG CGG CCC TCG CTG TGC CAG TTC ATC 109 Met Glu Ser Gly Gly Arg Pro Ser Leu Cys Gln Phe Ile -20 -15CTC CTG GGC ACC TCT GTG GTC ACC GCC GCC CTG TAC TCC GTG TAC 157

Leu Leu Gly Thr Thr Ser Val Val Thr Ala Ala Leu Tyr Ser Val Tyr

-10

### (2) INFORMATION FOR SEQ ID NO: 189:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 481 base pairs(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE
(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 133..355

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 3..225 id H10707

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 353..482

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 224..353

id H10707

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 154..354

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 93

region 98..298

id H30624

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 36S..403

(C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 94

region 314..349

id H30624 est

### (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 200..354

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 150..304 id HSC1VG011

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 111..198

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 92

region 62..149 id HSC1VG011

est

### (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 49..85

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 1..37 id HSC1VG011

### (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 202..344

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 113..255

id R34406

est

# (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 111..198

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 23..110

id R34406

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 353..482

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 115..244

id HSC23C111

(ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 240..355 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 94 region 1..116 id HSC23C111 (ix) FEATURE: (A) NAME/KEY: sig\_peptide (B) LOCATION: 56..472 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 4.3 seq GILVPHSLRQAQA/SF (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 189: AAAAACTGCG GAGGGTGACA AGGAAGAAGG TGGCTCCAGA TCTGGAGGTG TGTCC ATG 58 GCG GCG CTT GAC CTG CGA GCG GAS TGG ATT CGC TGG TCC TGC AGC TGC Ala Ala Leu Asp Leu Arg Ala Xaa Trp Ile Arg Trp Ser Cys Ser Cys -135 -130 TTG GGG GAM CTG GRA GGA GCT GGA GGG GAA ACG AAC GGT GTT GAA CGC 154 Leu Gly Xaa Leu Xaa Gly Ala Gly Gly Glu Thr Asn Gly Val Glu Arg -120 -115 -110 CCG GGT GGA GGG CTG GCT CTC GCT CGC CAA GGC TCG CTA CGC GAT 202 Pro Gly Gly Gly Leu Ala Leu Ala Arg Gln Gly Ser Leu Arg Asp -105 -100 GGG CGC CAA GTC GGT AGG GCC CCT GCA GTA TGC TTC CCA CAT GGA GCC Gly Arg Gln Val Gly Arg Ala Pro Ala Val Cys Phe Pro His Gly Ala -85 -80 CCA GGT CTG CCT CCA CGC CAG CGA GDC YCA GGA GGG DST CCA GAA GTT 298 Pro Gly Leu Pro Pro Arg Gla Arg Xaa Xaa Gly Gly Xaa Pro Glu Val CAA GGT GGT GAG AGC TGG TGT CCA CGC CCC AGA GGA GGT GGG GCC TCG 346 Gln Gly Gly Glu Ser Trp Cys Pro Arg Pro Arg Gly Gly Gly Ala Ser -50 CGA ACA GGT CTG CGG AGG CGC AAG GGC CCC ACT AAG ACC CCA GAA CCG 394 Arg Thr Gly Leu Arg Arg Arg Lys Gly Pro Thr Lys Thr Pro Glu Pro

-3.5

-20

GTT CCT CAC AGT CTA CGT CAG GCT CAA GCA AGC TTC CGG

Val Pro His Ser Leu Arg Gln Ala Gln Ala Ser Phe Arg

-5

-10

GAG TCC TCT GAG GCC CCT CAG GAC CCC CTG AAC TGG TTT GGA ATC CTA

Glu Ser Ser Glu Ala Pro Gln Asp Pro Leu Asn Trp Phe Gly Ile Leu

442

481

							103		
(2)	INFORMATION	FOR	SEQ	ΙD	NO:	190:			

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 302 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 176..275

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100 region 2..101

id R68368

est

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 216..278

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.1

seq WWISLLPSLLSIC/KV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 190:

AAGCTTTCCC CGTGGTCTGA GTTTGTGGCT GCATTTTTAT CTCTGGTGGC TCTGCTACGG 60

CGGCGCAGAA ATGAGGCAGA AGCGGAAAGG AGATCTCAGC CCTGCTGAGC TGATGATGCT 120

GACTATAGGA GATGTTATTA AACAACTGAT TGAAGCCCAC GAGCAGGGGA AAGACATCGA 180

TCTAAATAAG GTGAAAACCA AGACAGCTGC CAAAT ATG GCC TTT CTG CCC AGC Met Ala Phe Leu Pro Ser

-20

CCC GCC TGG TGG ATA TCA TTG CTG CCG TCC CTC AGT ATC TGC AAG 281 Pro Ala Trp Trp Ile Ser Leu Leu Pro Ser Leu Leu Ser Ile Cys Lys

GTC TTG ATG CCC AAG TTA AAG

302 Val Leu Met Pro Lys Leu Lys

### (2) INFORMATION FOR SEQ ID NO: 191:

(i) SEQUENCE CHARACTERISTICS: .

(A) LENGTH: 414 base pairs

(3) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii)	MOLECULE TYPE: CDNA	
(vi)	ORIGINAL SOURCE: (A) ORGANISM: Homo Sapid (F) TISSUE TYPE: Brain	ens
(ix)	FEATURE: (A) NAME/KEY: other (B) LOCATION: 40271 (C) IDENTIFICATION METHO (D) OTHER INFORMATION:	
(ix)	FEATURE: (A) NAME/KEY: other (B) LOCATION: 294328 (C) IDENTIFICATION METHOD OTHER INFORMATION:	
(ix)	FEATURE: (A) NAME/KEY: other (B) LOCATION: 86130 (C) IDENTIFICATION METH (D) OTHER INFORMATION:	
(ix)	FEATURE: (A) NAME/KEY: sig_pepti (B) LOCATION: 34180 (C) IDENTIFICATION METH (D) OTHER INFORMATION:	OD: Von Heijne matrix
(xi)	SEQUENCE DESCRIPTION: SE	Q ID NO: 191:
AATTCCCCAG	CAAGCTCAGC GTGTAMSTGC GC	T ATG GAG CCG AAA GTC GCA GAG 54 Met Glu Pro Lys Val Ala Glu -45
	n Lys Ile Glu Asp Thr Leu	TGT CCT TTT GGC TTC GAG GTT 102 Cys Pro Phe Gly Phe Glu Val -30
		GAA CTC TTG CCT CCA GCC TTC 150 Glu Leu Leu Pro Pro Ala Phe -15
		G GCC TTC CTG GTA CTC AGC ACG 198 1 Ala Phe Leu Val Leu Ser Thr 1 5
		G CCC TTC TTG CAG AGC TGC CAC 246 s Pro Phe Leu Gln Ser Cys His

10 15 20

CTC CGA ATG CTG ACT GAC CCA GTG GAC CAG TGT GTG GCC TAC CAT CTG
Leu Arg Met Leu Thr Asp Pro Val Asp Gln Cys Val Ala Tyr His Leu
25 30 35

GGC CGT GTT AGA GAG AGC CTC CCA GAG CTG CAG ATA GAA ATC ATT GCT

Gly Arg Val Arg Glu Ser Leu Pro Glu Leu Gln Ile Glu Ile Ile Ala

40

45

GRA HMA CGA GGT GCA CCC CAA CCG ACG CCC CAA GAT CCT GGC CCA GAC

Xaa Xaa Arg Gly Ala Pro Gln Pro Thr Pro Gln Asp Pro Gly Pro Asp

55 60 65 70

AGC AGC CAT GTA GCT GGG GCT GCT

Ser Ser His Val Ala Gly Ala Ala

75

### (2) INFORMATION FOR SEQ ID NO: 192:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 400 base pairs
- (B) TYPE: NUCLEIC ACID
- (C) STRANDEDNESS: DOUBLE
- (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens
(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 324..389

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95 region 301..366 id T08430

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 64..400

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 90

region 1..337 id C17891

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 301..400

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95 region 1..100

id C04989

WO 99/06552	168	PCT/IB98/01236
(iv) FEATURE:		

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 3.7

seq MLVLRSGLTKALA/SR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 192:

(A) NAME/KEY: sig\_peptide (B) LOCATION: 107..145

AGA(	STTCO	SCC A	AGTG	STCC	AG GA	AGCCO	CTT	TTT	CCAC	CTCG	GGA	AGACT	TTC A	AGAGA	AAGTCT	60
CAC	AAAGO	GAC 1	rcgg	CTGG	CT GO	CTTTT	CTC	A GTO	GCCG#	AAGC	CGC			CTC ( Leu \		115
	AGA Arg															163
	GTG Val															211
	TAT Tyr															259
	ATG Met 40															307
															AAA Lys 70	355
	TTT Phe															400

# (2) INFORMATION FOR SEQ ID NO: 193:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 186 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 112..184

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98 region 1..73 169

id HSC09D101 est

7:	· · ·	FEATURE
		FEATURE

(A) NAME/KEY: other (B) LOCATION: 112..184

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 1..73 id HSC2UE011

#### (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 112..186

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 90

region 1..75 id HSC09C101

est

### (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 140..184

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 17..61 id T35421 est

### (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 106..174

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 10.5

seq LLFVLLLFSLLPA/CL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 193:

ATATTTAAAC GACCCTTCAA AGGCCCTTAG GTTTCCTTGC CTCTGCTCAC AGAACTAGTC

CAGCCAGGTG TCGCTGCTGC CTCAGAGCTG TGTGGGGTCG CRTGT ATG TCG GGG GGC 117 Met Ser Gly Gly

-20

CAT CTT GCC GAT TTA ACG CTG CTT TTT GTG TTG TTG TTT TCC CTC 165 His Leu Ala Asp Leu Thr Leu Leu Phe Val Leu Leu Leu Phe Ser Leu

-15

CTC CCT GCC TGC CTA CCC CGG Leu Pro Ala Cys Leu Pro Arg 186

# (2) INFORMATION FOR SEQ ID NO: 194:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 335 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR

#### (ii) MOLECULE TYPE: CDNA

### (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement(86..336)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 79..329 id AA148596 est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (30..91)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 325..386 id AA148596

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (2..39)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 92

region 378..415 id AA148596

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement(2..336)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 67..401 id AA074631

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement(83..336)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 64..317 id AA078818

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement(8..48)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 355..395 id AA078818

		est
(ix)	FEATURE:  (A) NAME/KEY: other  (B) LOCATION: complement  (C) IDENTIFICATION METHO  (D) OTHER INFORMATION:	
(ix)	FEATURE:  (A) NAME/KEY: other  (B) LOCATION: complement  (C) IDENTIFICATION METHO  (D) OTHER INFORMATION:	
(ix)	FEATURE:  (A) NAME/KEY: other  (B) LOCATION: complement  (C) IDENTIFICATION METHO  (D) OTHER INFORMATION:	DD: blastn
(ix)	FEATURE:  (A) NAME/KEY: other  (B) LOCATION: complement  (C) IDENTIFICATION METHO  (D) OTHER INFORMATION:	
(ix)	FEATURE:  (A) NAME/KEY: sig_peptic (B) LOCATION: 174326 (C) IDENTIFICATION METHO (D) OTHER INFORMATION:	
(xi)	SEQUENCE DESCRIPTION: SE	Q ID NO: 194:
AATTCTCAAC	GAGCTGCGGG CTCGGCATGC CC	AGGGGGGT ACATGGTATG GAGTAGACAT 60
CAACAACGAG	GACATTGCTG ACAACTTTGA AG	CTTTCGTG TGGGAGCCAG CTATGGTGCG 120
GATCAATGCG	CTGACAGCAG CCTCTGAGGC TG	CGTGCCTG ATCGTGTCTG TAG ATG 176 Met

AAA CCA TCA AGA ACC CCC GCT CGA CTG TGG ATG CTC CCA CAG CAG CAG

Lys Pro Ser Arg Thr Pro Ala Arg Leu Trp Met Leu Pro Gln Gln

GCC GGG GCC GTG GTC GTG GCC GCC CCC ACT GAG AGG CAC CCC ACC CAT

Ala Gly Ala Val Val Ala Ala Pro Thr Glu Arg His Pro Thr His

-40

-45

224

272

-25 -30

CAC ATG GCT GGC TGG CTG GGT GCA CTT ACC CTC CTT GGC TTG GTT 320 His Met Ala Gly Trp Leu Leu Gly Ala Leu Thr Leu Leu Gly Leu Val

-10 -15

ACT TCA TTT TAC AAG Thr Ser Phe Tyr Lys

335

-20

# (2) INFORMATION FOR SEQ ID NO: 195:

### (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 419 base pairs
- (B) TYPE: NUCLEIC ACID
- (C) STRANDEDNESS: DOUBLE
- (D) TOPOLOGY: LINEAR

### (ii) MOLECULE TYPE: CDNA

### (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo Sapiens
- (F) TISSUE TYPE: Brain

### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 23..200
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 100 region 9..186

id W44639

# (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 216..356
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 99

region 200..340

id W44639

est

### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 363..412
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 100

region 347..396

id W44639

# (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 27..218
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 9.5

seq LSLLAALAHLAAA/EK

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 195:

AGTO	GGTC	GA H	CTG	GGCC	GC AC	STCGO			Pro		GCC Ala	53
	GTC Val											101
	ATC Ile											149
	GCC Ala											197
	GCG Ala											245
	GGG Gly											293
	CGC Arg											341
	GGC Gly											389
	TTC Phe											419

# (2) INFORMATION FOR SEQ ID NO: 196:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 342 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA-
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 33..269
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 99 region 10..246 id AA058587

est

(ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 272..307

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 248..283 id AA058587

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 133..259

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 87..213 id R12128

20 412120

(ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 47..134

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 2..89 id R12128

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 303..337

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 257..291

id R12128

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 49..259

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 2..212

id H19999

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 272..304

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 225..257

id #19999

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 303..337

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 91

WO 99/06552 PCT/IB98/01236

region 257..291 id H19999

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 42..252

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 1..211 id R20025 est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 87..259

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 1..173 id H83838 est

(ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 272..337

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 186..251 id H83838

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 85..198

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 9.3

seq QLLYLSLLSGLHG/QE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 196:

AGGCTGCGGT AAATCCGGGC TTGCGGCCGC TGGCGTAGTC TGTGGCCGGG TGGTCGTTGC 60

TGCGCGCCCC GAGCCCCGAG AGCC ATG CAG ATG TCC TAC GCC ATC CGG TGC

Met Gln Met Ser Tyr Ala Ile Arg Cys

-35

-30

GCC TTC TAC CAG CTG CTG GCC GCG CTC ATG CTG GCG ATG CTG
Ala Phe Tyr Gln Leu Leu Ala Ala Leu Met Leu Val Ala Met Leu
-25
-20
-15

CAG CTG CTC TAC CTG TCG CTG CTG TCC GGA CTG CAC GGG CAG GAG GAG GIn Leu Leu Tyr Leu Ser Leu Leu Ser Gly Leu His Gly Gln Glu

Tyr Leu Ser Leu Leu Ser Gly Leu His Gly Gln Glu -10 -5 1

CAA GAC CAA TAT TTT GAG TTC TTT CCC CCG TCC CCA CGG TCC GTG GAC 255
Gln Asp Gln Tyr Phe Glu Phe Phe Pro Pro Ser Pro Arg Ser Val Asp
5 10

CAG GTC AAG GCT CAG CTC CGC ACC GCG CTG GCC TCT GGA GGC GTC CTG 303 Gln Val Lys Ala Gln Leu Arg Thr Ala Leu Ala Ser Gly Gly Val Leu

PCT/IB98/01236 WO 99/06552 176

25 30 35

GAC GCT AGC GGC GAT TAC CGC GTC TAC AGG GGC CAT GGG Asp Ala Ser Gly Asp Tyr Arg Val Tyr Arg Gly His Gly 40 45

342

# (2) INFORMATION FOR SEQ ID NO: 197:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 461 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: complement(149..337)
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 97 region 182..370 id AA142966

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: complement (340..459)
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 97 region 61..180 id AA142966 est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: complement (142..337)
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 98 region 183..378

id AA019334

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: complement (340..459)
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 97 region 62..181

id AA019334

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: complement (345..459)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 48..162

id N66447

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (255..337)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 170..252

id N66447

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement(111..181)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 330..400

id N66447

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (179..228)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 282..331

id N66447

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement(172..337)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97 region 113..278

id R85770

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (340..450)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 1..111

id R85770

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 188..337

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 1..150

id R78830

est

### (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 339..459

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 151..271 id R78830

00

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 384..455

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 9.1

seq LFAFHLLLSFILG/SR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 197:

AACACTGTTG TATAAACTAA TCTTTGCTTG TTTTCTACTC TGTGATCTTT CCATATCATA 60 TTTCATTAAT GATCAGTTAG TGTCAAGGAG TCAAAACAGA TTAAAATTAA TTTCATGTGT 120 ATATGGTGGA AATTTGTGGC TAGTGTGATT TTTGTTTGTY TCCTTTTAAG TACTGTTGAT 180 CAGTTGTGAC ACTTACTGGT TAAACTTACG TTGCTAAAGA TTTCTCTATA ATAAGCCACA CATTATATT AGACTATATT AAGGGACCTT GGTTTTCTTC TAGATAGCAG CTGTCCCAAA 300 GAAAATATTT CTTCTTTGTC TGTKAAGATT TAGCTATNKA TCTGCCAGTT GTTCAGMGGT 360 TTTGGTTCCA AACTCAACCA GCA ATG TTG AGA GCT GAA CTT AAG ATA GCT GTT 413 Met Leu Arg Ala Glu Leu Lys Ile Ala Val GTA CTT TTT GCT TTC CAT CTG TTA CTG TCC TTC ATT CTC GGC TCC CGG 461 Val Leu Phe Ala Phe His Leu Leu Leu Ser Phe Ile Leu Gly Ser Arg -10 -5

# (2) INFORMATION FOR SEQ ID NO: 198:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 229 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: complement(1..130)
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 98 region 8..137 id H63707

est

(ix)	FEATURE:
	(A) NAME/KEY: sig_peptide
	(B) LOCATION: 2166
	(C) IDENTIFICATION METHOD: Von Heijne matri:
	(D) OTHER INFORMATION: score 9
	seq LLILLLRTFLCSA/M
(xi)	SEQUENCE DESCRIPTION: SEQ ID NO: 198:

A ATG AAT CAT CAA CAG ACA TTA ATT GGG CGC CTG CTC TGT GAT CTC CAT Met Asn His Gln Gln Thr Leu Ile Gly Arg Leu Leu Cys Asp Leu His -50 -45 GGG CTC AGC TTG TCC CCG CCA GTW GCC AAC AAC GTC CAA GCT CTC TTC 97 Gly Leu Ser Leu Ser Pro Pro Val Ala Asn Asn Val Gln Ala Leu Phe -35 AGA ATG CTT ACT CCT GAA GCT TAT TCC TGT CTT CTA ATT CTT TTG TTG 145 Arg Met Leu Thr Pro Glu Ala Tyr Ser Cys Leu Leu Ile Leu Leu -20 -15 AGG ACT TTT CTG TGT AGT GCA ATG ATA GCA AAT ACA CTT CAT CTC AAG 193 Arg Thr Phe Leu Cys Ser Ala Met Ile Ala Asn Thr Leu His Leu Lys 1 TAC CAT CTC CAA TTG ATT GAT AAT GCC TGC CCT GAG 229 Tyr His Leu Gln Leu Ile Asp Asn Ala Cys Pro Glu 10 15

# (2) INFORMATION FOR SEQ ID NO: 199:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 278 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 145..279
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 100 region 1..135 id H06014

- (ix) FEATURE:
  - (A) NAME/KEY: other (B) LOCATION: 180..279
  - (C) IDENTIFICATION METHOD: blastn

. (D) OTHER INFORMATION: identity 100 region 11..110 id R15960

est

lix	) FFA	TURE

(A) NAME/KEY: other (B) LOCATION: 204..279

(C) IDENTIFICATION METHOD: blastn

. (D) OTHER INFORMATION: identity 96

region 1..76 id W67034 est

# (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 27..146

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 8.6

seq LFCVLGIVLLVTG/IV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 199:

ACCAAACCAA AGCCGTCATC ATTGCA ATG ATC ATC ACT GCG GTG GTA TCC ATT

Met Ile Ile Thr Ala Val Val Ser Ile

-40

-35

TCA GTC ACC ATC TTC TGC TTT CAG ACC AAG GTG GAC TTC ACC TCG TGC

Ser Val Thr Ile Phe Cys Phe Gln Thr Lys Val Asp Phe Thr Ser Cys

-30

-25

-20

ACA GGC CTC TTC TGT GTC CTG GGA ATT GTG CTC CTG GTG ACT GGG ATT

Thr Gly Leu Phe Cys Val Leu Gly Ile Val Leu Leu Val Thr Gly Ile
-15
-10
-5
1

GTC ACT AGC ATT GTG CTC TAC TTC CAA TAC GTT TAC TGG CTC CAC ATG

Val Thr Ser Ile Val Leu Tyr Phe Gln Tyr Val Tyr Trp Leu His Met

5 10 15

CTC TAT GCT GCT CTG GGG GCC ATT TGT TTC ACC CTG TTC CTG GCT TAC
Leu Tyr Ala Ala Leu Gly Ala Ile Cys Phe Thr Leu Phe Leu Ala Tyr
20 25 30

GAC ACA CAG CTG GTC CTG GGG AAC CGG AAG CAC
Asp Thr Gln Leu Val Leu Gly Asn Arg Lys His
35 40

### (2) INFORMATION FOR SEQ ID NO: 200:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 333 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE
(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 55..268 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 95 region 32..245 id T60555 (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 22..51 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 90 region 1..30 id T60555 est (ix) FEATURE: (A) NAME/KEY: sig\_peptide (B) LOCATION: 67..261 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 8.4 seq LLWFIHLVFVVLX/LF (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 200: AAGAGGCTTA CGAGSWCCAG GTGGAGAGGC CGGGCTGGCC AAGGCTTCGG CCTCCGGCGT 60 CGGGAA ATG GCG GCG GGG GGC AGG ATG GAG GAC GGT TCC TTG GAT ATC 108 Met Ala Ala Gly Gly Arg Met Glu Asp Gly Ser Leu Asp Ile -60 ACC CAG AGT ATT GAA GAC GAC CCA CTT CTG GAT GCC CAG CTT CTC CCA Thr Gln Ser Ile Glu Asp Asp Pro Leu Leu Asp Ala Gln Leu Leu Pro -45 CAC CAC TCA TTA CAA GCT CAC TTT AGA CCC CGA TTC CAT CCT CTT CCT 204 His His Ser Leu Gln Ala His Phe Arg Pro Arg Phe His Pro Leu Pro ACA GTC ATC ATA GTG AAT CTT CTG TGG TTT ATT CAT CTC GTG TTT GTT Thr Val Ile Ile Val Asn Leu Leu Trp Phe Ile His Leu Val Phe Val -15 -10 GTW TTA GSA TTG TTT AAC AGG TGT GCT TTG TTC TWA TCC TAT CCC AAA 300 Val Leu Xaa Leu Phe Asn Arg Cys Ala Leu Phe Xaa Ser Tyr Pro Lys TGG GAC ARG TGC CCA GGA AAT TAC ACA AAC CCA 333 Trp Asp Xaa Cys Pro Gly Asn Tyr Thr Asn Pro

(2) INFORMATION FOR SEQ ID NO: 201:

20

15

(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 337 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPÓLOGY: LINEAR (ii) MOLECULE TYPE: CDNA (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 125..306 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 95 region 95..276 id H31193 est (ix) FEATURE: (A) NAME/KEY: other (3) LOCATION: 69..130 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 93 region 40..101 id H31193 (ix) FEATURE: (A) NAME/KEY: other (3) LOCATION: 29..68 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 97 region 1..40 id H31193 (ix) FEATURE: (A) NAME/KEY: sig\_peptide (B) LOCATION: 161..208 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 7.5 seq GCMLLFVFGFVGG/AV (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 201: AATCGCTTGG GAGCTGCTGC AGGATGGAGT GGAAAGCTGC TGCTGATGGC ATTGTTTTTG TGGCAGCAAG CTGAATGACA GATCCTCACT ACAAAGATAC CCCTTTGGCC CCCGTGTAGG 120 CCTCCTTGGT TCGGGTGTTT CACCATGCCA GCACAGCGCC ATG AGT CCT GGA TGC Met Ser Pro Gly Cys -15 223 Met Leu Leu Phe Val Phe Gly Phe Val Gly Gly Ala Val Ile Asn

TCT GCT ATC TTA GTA TCT CTC TCT GTT TTG CTG CTT GTG CAC TTT TCT Ser Ala Ile Leu Val Ser Leu Ser Val Leu Leu Leu Val His Phe Ser 10

15

ATT TCT ACC GGT GTG CCA GCT CTG ACG CAG AAC CTA CCA AGG ATA CTC 319 Ile Ser Thr Gly Val Pro Ala Leu Thr Gln Asn Leu Pro Arg Ile Leu 30

AGA AAA GAA CGC CCC GGG Arg Lys Glu Arg Pro Gly 40

337

## (2) INFORMATION FOR SEQ ID NO: 202:

# (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 309 base pairs
- (B) TYPE: NUCLEIC ACID
- (C) STRANDEDNESS: DOUBLE
- (D) TOPOLOGY: LINEAR

## (ii) MOLECULE TYPE: CDNA

## (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo Sapiens
- (F) TISSUE TYPE: Brain

### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 105..252
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 94

region 136..283 id HSU46355

est

# (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 53..83
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 90

region 82..112

id HSU46355 est

# (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 227..276
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 94

region 206..255

id AA011705

est .

# (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 109..153
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 7.5

seq LLLGIALLAYVAS/VW

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 202:

AATBGTGCAG CAGGCGGGCC CCCGCGCGGC AGGGSCCTGG ACCCGCGCGG CTCCCTGGGA 60 TGGTGAGCAA GGCGCTGCTG CSCTCGTGTC TGCCGTCAAC CGCAGASG ATG AAG CTG 117 Met Lys Leu -15 CTG CTG GGC ATC GCC TTG CTG GCC TAC GTC GCC TCT GTT TGG GGC AAC 165 Leu Leu Gly Ile Ala Leu Leu Ala Tyr Val Ala Ser Val Trp Gly Asn -10 -5 TTC GTT AAT ATG AGG TCT ATC CAG GAA AAT GGT GAA CTA AAA ATT GAA 213 Phe Val Asn Met Arg Ser Ile Gln Glu Asn Gly Glu Leu Lys Ile Glu 10 AGC AAG ATT GAA GAG ATG GTT GAA CCA CTA AGA GAG AAA ATC AGA GAT 261 Ser Lys Ile Glu Glu Met Val Glu Pro Leu Arg Glu Lys Ile Arg Asp 25 TTA GRA AAA AGC TTT ACC CAG AAA TAC CCA CCA GTA AAG TTT TTA TCA 309 Leu Xaa Lys Ser Phe Thr Gln Lys Tyr Pro Pro Val Lys Phe Leu Ser 40

### (2) INFORMATION FOR SEQ ID NO: 203:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 491 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 132:.251
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 99

region 170..289

id T60981

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 19..126
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 99

region 57..164

id T60981

est

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 39..107

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 7.5

seq LVLLLTLPLHLMA/LL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 203:

AAGTGCCCCA GCGGAAGACA GCTCAGAGCT GGTCTGCC ATG GAC ATC CTG GTC CCA Met Asp Ile Leu Val Pro -20														56	
					GTG Val										104
					TGG Trp 5						Ser				152
					ACT Thr										200
					AGC Ser										248
					GAG Glu										296
					TGC Cys										344
					ACA Thr 85									 	392
					GTG Val										440
					GTK Val										488
CAG Gln															491

# (2) INFORMATION FOR SEQ ID NO: 204:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 331 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

PCT/IB98/01236 WO 99/06552 186

(ii) MOLECULE TYPE: CDNA

#### (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 25..303

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 1..279 id HSC0ZA041

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 131..286

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 106..261

id R12615

est

### (im) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 71..133

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 47..109

id R12615

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 88..303

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99 region 1..216

id HUM401H04B

## (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 137..303

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95 region 92..258

id T78771

est

## (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 23..127

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 7.4

seq SLLLSLELASGSG/QG

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 204:

AAAGGGCGGA STTCAGGTCT CC ATG GAG GCG GCT TCT CCT AGC AAC TCG ACG 52 Met Glu Ala Ala Ser Pro Ser Asn Ser Thr -35GGC GTT GAG CGG ASC GCT GAC CTG ATG GAC GCC GAC AGC CTC CTG CTG 100 Gly Val Glu Arg Xaa Ala Asp Leu Met Asp Ala Asp Ser Leu Leu Leu -25 TCT CTG GAG CTG GCG TCC GGC AGT GGG CAG GGC CTC AGC CCG GAC CGT 148 Ser Leu Glu Leu Ala Ser Gly Ser Gly Gln Gly Leu Ser Pro Asp Arg CGG GCC TCG CTC ACG TCT CTT ATG CTG GTT AAG CGC GAC TAC CGC 196 Arg Ala Ser Leu Leu Thr Ser Leu Met Leu Val Lys Arg Asp Tyr Arg 15 TAT GAT CGG GTT CTC TTC TGG GGC CGC ATC CTT GGC CTC GTC GCC GAT 244 Tyr Asp Arg Val Leu Phe Trp Gly Arg Ile Leu Gly Leu Val Ala Asp 30 TAC TAC ATC GCG CAG GGC CTG AGT GAG GAC CAG CTC GCA CCG CGC AAG 292 Tyr Tyr Ile Ala Gln Gly Leu Ser Glu Asp Gln Leu Ala Pro Arg Lys 45 ACG CTC TAT AGG TCC AGA TCA AGG AAG AGA CCC GCA CTG 331 Thr Leu Tyr Arg Ser Arg Ser Arg Lys Arg Pro Ala Leu 60 65

## (2) INFORMATION FOR SEQ ID NO: 205:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 317 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 46..119
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 93

region 12..85

id N80892

id H92328

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 88..119
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 100 region 1...32

est

(	ix)	FEATURE	:
---	-----	---------	---

- (A) NAME/KEY: sig\_peptide
  (B) LOCATION: 108..236
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 7.4

seq VLVKLLSSSASTS/RP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 205:

AGTTTCGNST CGCGGATCCG GTAGGTCCAG GTGCAGCGGC CGCAGTKCTG CGTCCGTGCG 60														
CCG	CGGGC	CTG (	GGCC	GTC	rc ac	GTGT	rgcco	AAC	GCTC1	rggt	CAG	rgcc	 ATC Ile	 116
	GAG Glu												 	 164
	AGC Ser													212
	TCC Ser													260
	CAA Gln 10													 308
	CCA Pro													317

# (2) INFORMATION FOR SEQ ID NO: 206:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 363 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 26..365
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 96 region 1..340

id N40260

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 17..308

(C) IOENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 93

region 6..297 id W07706 est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 311..349

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 301..339 id W07706

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 79..365

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 22..308 id W37568

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 140..326

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 93

region 74..260

id W00732

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 328..365

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 263..300

id W00732

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 79..362

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 14..297

id AA135041

est

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 25..147

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 7.4

seq ILPLLFGCLGVFG/LF

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 206:

ACACGTCACT TCCGAGGCGG GAGG ATG AAG TTG ATT GAC TAT GGT CTC TCC  Met Lys Leu Ile Asp Tyr Gly Leu Ser  -40 -35													51		
					TCT Ser										99
					CCC Pro										147
					CAG Gln										195
					ACA Thr										243
					GCT Ala								 		291
					GAA Glu										339
					ACA Thr 70										363

# (2) INFORMATION FOR SEQ ID NO: 207:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 235 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 60..181
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 97 region 1..122 id AA057454

(ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 182..233

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 122..173 id AA057454

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 71..233

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 1..163 id C18312 est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 182..233

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 144..195

id W69247

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 98..144

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 62..108

id W69247

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 34..78

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 1..45

id W69247

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 146..233

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 69..156

id H75891

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 76..144

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 1..69

	est	
(B) (C)	URE:  NAME/KEY: other  LOCATION: 80233  IDENTIFICATION METHOD: blastn  OTHER INFORMATION: identity 98  region 1154  id HUML11265  est	
(B) (C)	URE: NAME/KEY: sig_peptide LOCATION: 104160 IDENTIFICATION METHOD: Von Heijne matrix OTHER INFORMATION: score 7.4 seq PMLLRALAQAARA/GP	
(xi) SEQU	ENCE DESCRIPTION: SEQ ID NO: 207:	
ATAAGGGGGA ACCC	GCTGGC CCAATGGCAG CGTCCTACAG TGTAGCCTCC GCCTCCCGAT	60
TGACTGGCCT GCTT	GGCAAK GCAAGTAGCG GCGGCGCTTC AAG ATG CGC TGC CTG Met Arg Cys Leu	115
	CTG CTG CGG GCC CTG GCC CAG GCT GCA CGT GCA GGA Leu Leu Arg Ala Leu Ala Gln Ala Ala Arg Ala Gly -10 -5 1	163
	CGG AGC CTC CAC AGC AGT GCA GTG GCA GCC ACC TAC Arg Ser Leu His Ser Ser Ala Val Ala Ala Thr Tyr 10	211
	ATG CAG GAT CAA Met Gln Asp Gln 25	235
(2) INFORMATION	FOR SEQ ID NO: 208:	
(A) (B) (C)	NCE CHARACTERISTICS: LENGTH: 385 base pairs TYPE: NUCLEIC ACID STRANDEDNESS: DOUBLE TOPOLOGY: LINEAR	
(ii) MOLE	CULE TYPE: CDNA	
(A)	INAL SOURCE: ORGANISM: Homo Sapiens TISSUE TYPE: Brain	
(B) (C)	NAME/KEY: other LOCATION: 70351 IDENTIFICATION METHOD: blastn OTHER INFORMATION: identity 97 region 34315 id T19063	

est

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(ix) FEATURE:
```

(A) NAME/KEY: other (B) L@CATION: 36..68

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 1..33 id T19063 est

## (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 61..353

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 1..293 id T32338 est

## (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 93..360

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 1..268 id T30463 est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (107..265)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 91

region 330..488 id W27204

est

## (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (257..385)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 209..337

id W27204

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 70..324

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 27..281

id T32187

est

### (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 134..334

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 7

# seq IWTLLSSVIRCLC/AI

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 208:

AGACAAA'	rgg Ci	CAGGTG	GA CTCCG	GGCTG GA	GCTGTCCT	GGGGGAGC	TT GTTT	GCGGCA	60
SGGCTGC:	rgc to	GCCACTG	CT GTGCT	ggsgg co	CGGTCGCC	AGGCAAAA	AG CCCT	CCCACG	120
TTTGAGG	GGA GI				Asn Val	TTA AGA A Leu Arg S -60			169
						CTG CAG Leu Gln			217
						GGC AAG Gly Lys			265
	Gly I				Gly Ile	TGG ACG Trp Thr			313
						CAC AAC His Asn 5			361
			TGG ACC Trp Thr 15						385

# (2) INFORMATION FOR SEQ ID NO: 209:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 285 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: complement(2..55)
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 92 region 34..87

id T86932

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: complement (45..86)

(C) IDENTIFICATION METHOD: blastn
(D) OTHER INFORMATION: identity 100
region 2..43

id T86932

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 199..240

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 6.8

seq IFLTLSLDSRVSA/IR

285

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 209:

AAATAAAAAT ATCTTAAAAC TGCATTGTAC AGCTCCCTCC CTGCGTTTTA TTAAATGATG 60
TATATTAAAC AAAGATCAAT ATTTTCTTAA TGACTCAGGG TCTTTATTGT TAATGCCAAT 120

TGTTTTTGTA TCTGTGCTAT AATCCCTTAG AGTCAGTAAA GTATGTAGGG GACTGTTTCT 180

TCCTTTGTGT CTGGGTTT ATG ATT TTT CTC ACT CTT TCT TTG GAC TCC AGG

Met Ile Phe Leu Thr Leu Ser Leu Asp Ser Arg

-10

-5

GTG TCA GCC ATC AGG TCT CCT AAT TTT GTG TAC CGG TCT CCA ACA DMC 279
Val Ser Ala Ile Arg Ser Pro Asn Phe Val Tyr Arg Ser Pro Thr Xaa

1 5 10

CAT GGG His Gly 15

(2) INFORMATION FOR SEQ ID NO: 210:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 378 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 65..270

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97 region 109..314

id AA100852

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 269..378

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 314..423 id AA100852

est

## (ix) FEATURE:

- (A) NAME/KEY: other (B) LOCATION: 65..270
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 97

region 109..314 id AA161042

est

# (ix) FEATURE:

- (A) NAME/KEY: other(B) LOCATION: 277..361
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 94

region 323..407 id AA161042

est

### (ix) FEATURE:

- (A) NAME/KEY: other(B) LOCATION: 65..274
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 97

region 104..313 id H64488

est

### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 68..256
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 97

region 147..335 id AA146605

est

### (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 256..317

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 336..397 id AA146605

est

### (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 80..305

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 129..354

id AA088770

<pre>(ix) FEATURE:     (A) NAME/KEY: sig_peptide     (B) LOCATION: 76162     (C) IDENTIFICATION METHOD: Von Heijne matrix     (D) OTHER INFORMATION: score 6.8</pre>																
	(:	xi) S	SEQUI	ENCE	DESC	CRIP	rion:	: SE	Q ID	NO:	210	:				
AAT	ACTA	CAC A	ACTC	ATAT	AG G	GGAG	GGAG	G CT	TCTG	GTC	CCA	GGC	CGC A	AGGG	GCAKKG	60
AAG'	rctg(	GAG (	CCWY						r Phe					t Me	G ATC	111
		AAT Asn -15														159
		TGG Trp														207
		TCG Ser														255
		GCC Ala														303
		AAG Lys 50														351
		CTC Leu														378
(2)	INF	ORMA'	TION	FOR	SEQ	ID	NO:	211:								
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 327 base pairs  (B) TYPE: NUCLEIC ACID  (C) STRANDEDNESS: DOUBLE  (D) TOPOLOGY: LINEAR																
	(.	ii)	MOLE	CULE	TYP	E: C	DNA									
	(	vi) (	(A)	ORG	ANIS	RCE: M: Ho TYPE			ens							
	(	ix)	(A)	NAM		Y: 0'		288								

(C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 96

region 203..257 id R25833 est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 285..317

(C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 100

region 255..287

id R25833

est

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 37..141

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 6.4

seq SACLLLCPTWTNP/QL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 211:

AAA	AAAGO	SCG (	GGT	CTCG	SC CC	GCGC	CTGAC	GC#	AGCC	 	 GCG Ala	 	54
							GAG Glu			 	 	 	102
							CCT Pro						150
							TCT Ser						198
							CGG Arg						246
							TCC Ser			 _	 	 	294
							GTD Val						327

# (2) INFORMATION FOR SEQ ID NO: 212:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 244 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 82..241

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 51..210 id C18780

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 48..83

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 18..53 id C18780

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 163..235

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 121..193 id T11911

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 116..162

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 73..119

id T11911

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 204..239

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 226..261

id T69629

est

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide
(B) LOCATION: 143..199

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 6.4

seq SVFLLMVNGQVES/AQ

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 212:

200 AGCACTCGCG TGGCCTTCGC GAAGGTGTCG CTGCCAAGAA ACGTGTCCTG CGCGCTACGC CGTCTGTTTT TAGGGCAACG CCGGCGTCTC TTAGCAACCG CGCGCGGCCT AGGTGGGTCC 120 CCCCGGCACC CCCAGACCTG CC ATG GCG ACC GCG AGT CCT AGC GTC TTT CTA Met Ala Thr Ala Ser Pro Ser Val Phe Leu -15 CTC ATG GTC AAC GGG CAG GTG GAG AGC GCC CAG TTT CCA GAG TAT GAT 220 Leu Met Val Asn Gly Gln Val Glu Ser Ala Gln Phe Pro Glu Tyr Asp GAC CTC TAC TGC AAG TAC TGC CAG 244 Asp Leu Tyr Cys Lys Tyr Cys Gln 10 (2) INFORMATION FOR SEQ ID NO: 213: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 211 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR (ii) MOLECULE TYPE: CDNA (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 95..208 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 100 region 85..198 id N43024 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 28..95 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 91 region 17..84 id N43024 est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 107..199

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98 region 80..172

id T62095 est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 61..106 (C) IDENTIFICATION METHOD: blastn (D) OTHER 1HFORMATION: identity 93 region 35..80 id T62095 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 26..60 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 97 region 1..35 id T62095 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 61..208 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 98 region 26..173 id W42796 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 110..208 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 92 region 114..212 id AA030227 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 110..208 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 92 region 51..149 id AA118270 (ix) FEATURE: (A) NAME/KEY: sig\_peptide (B) LOCATION: 104..187 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 6 seq IGLMFLMLGCALP/IY (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 213: TCTTCCGGGT GTTGTCTGGC CGCCGTAGCG CRTCTTGGGT CTCCCGGCTG CCGCTGCTGC CGCCGCCGCC TCGGGTCGTG GAGCCAGGAG CGACGTCACC GCC ATG GCA GGC ATC Met Ala Gly Ile AAA GCT TTG ATT AGT TTG TCC TTT GGA GGA GCA ATC GGA CTG ATG TTT 163 Lys Ala Leu Ile Ser Leu Ser Phe Gly Gly Ala Ile Gly Leu Met Phe

WO 99/06552 PCT/IB98/01236

-20 -15 -10

TTG ATG CTT GGA TGT GCC CTT CCA ATA TAC AAC AAA TAC TGG CCC TGG
Leu Met Leu Gly Cys Ala Leu Pro Ile Tyr Asn Lys Tyr Trp Pro Trp

-5

1

5

# (2) INFORMATION FOR SEQ ID NO: 214:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 128 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 3..124
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 99

region 8..129 id AA146587

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 2..124
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 98

region 14..136

id T85006

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 11..124
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 100

region 1..114

id H08511

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 14..124
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 98

region 1..111

id C00740

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 13..124

203 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 98 region 1..112 id N40664 (ix) FEATURE: (A) NAME/KEY: sig\_peptide (B) LOCATION: 12..62 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 5.9 seq ILLFGTLLMNAGA/VL (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 214: AGGCCGTAAC G ATG ATC GGA GAC ATC CTG CTG TTC GGG ACG TTG CTG ATG 50 Met Ile Gly Asp Ile Leu Leu Phe Gly Thr Leu Leu Met -10 AAT GCC GGG GCG GTG CTG AAC TTT AAG CTG AAA AAG AAG GAC ACG CAG 98 Asn Ala Gly Ala Val Leu Asn Phe Lys Leu Lys Lys Lys Asp Thr Gln 1 5 GGC TTT GGG GAG GAG TCC AGG GAG CCT TGG 128 Gly Phe Gly Glu Glu Ser Arg Glu Pro Trp 15 (2) INFORMATION FOR SEQ ID NO: 215: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 150 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR (ii) MOLECULE TYPE: CDNA (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain (A) NAME/KEY: other (B) LOCATION: 12..143 (C) IDENTIFICATION METHOD: blastn

#### (ix) FEATURE:

(D) OTHER INFORMATION: identity 99

region 36..167 id HUM137D01B

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 12..142

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99 region 143..273

id AA155928

(ix) FEATURE:

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

	E/KEY: other ATION: 12141		
(C) IDE	NTIFICATION METHO	OD: blastn identity 99 region 115244 id W39572 est	
(ix) FEATURE:	E/VEV. othor		
(B) LOC.	E/KEY: other ATION: complement NTIFICATION METHO ER INFORMATION:	·	
(ix) FEATURE:			
• •	E/KEY: other ATION: complement	t (32151)	
	NTIFICATION METH ER INFORMATION:		
(2, 0	J. 2.11. Giuli 12011	region 346465 id H99266 est	
(ix) FEATURE:			
(B) LOC. (C) IDE	E/KEY: sig_peption ATION: 67114 NTIFICATION METHOR ER INFORMATION:	OD: Von Heijne matrix	
(xi) SEQUENCE	DESCRIPTION: SE	Q ID NO: 215:	
ACACATCCCT CTAAACTA	CT GTTAGGAACA GC	AGTGTTCT CACAGTGTRG GGCAGCC	CGTC 60
		CTG TCC CTC TTT GGC AGT TGC Leu Ser Leu Phe Gly Ser Cys -5	
		GAG CGT AGC ATC CAG Glu Arg Ser Ile Gln 10	150
(2) INFORMATION FOR	SEQ ID NO: 216:		
(A) LEN (B) TYP (C) STR	CHARACTERISTICS: GTH: 397 base pa E: NUCLEIC ACID ANDEDNESS: DOUBL OLOGY: LINEAR		

(A) ORGANISM: Homo Sapiens
(F) TISSUE TYPE: Brain

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement(223..398)(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 111..286 id HSGT545 est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement(69..219)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 92

region 291..441 id HSGT545

est

### (ix) FEATURE:

(A) NAME/KEY: other

(3) LOCATION: complement (2..43)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 467..508 id HSGT545

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (223..311)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 92

region 4..92 id AA036134

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (46..163)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 91

region 133..250 id AA038839

est

## (ix) FEATURE:

(A) NAME/KEY: other

(3) LOCATION: complement(223..295)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 1..73 id AA038839

---

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 326..387

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 91 region 2..63 id W51392 est

# (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
  (B) LOCATION: 152..268
- (C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 5.9

seq SVSVLSSLGIVLA/VV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 216:

ACT	TGAC	GGG	TGTC	CTGC	SC C	ATGTO	GTGT	TTO	GATG	CCAG	CBG	CTCT	CGG	ATGG	CATGG	A 60
CGC1	TAT	CGA	GCAG	CTTCA	AG GO	STGG	CAGC	AC	AAGA	AGAT	TGG	CTAC'	TAT	GACA	CACC	A 120
AGG	ATGAT	гст	TTCC	rggto	CC A	AAAC <i>i</i>	AGAT <i>i</i>					Arg '		CCC (		172
			GAC Asp													220
			TCC Ser													268
			CTG Leu													316
			CAG Gln 20													364
			GCT Ala		Val	Phe										397

# (2) INFORMATION FOR SEQ ID NO: 217:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 373 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other

(B) LOCATION: 41..337(C) IDENTIFICATION METHOD: blastn(D) OTHER INFORMATION: identity 97

region 1..297 id H56523 est

## (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 38..337

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 1..300 id AA020823 est

## (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 43..337

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 7..301 id H99096 est

### (ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 49..315

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 11..277 id AA083141 est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 52..337

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 17..302 id N21197 est

## (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 35..82

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 5.8

seq AALPAWLSLQSRA/RT

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 217:

AGCTTGTCCC CTCCGGCTTG CCGTCCTCGC AGCC ATG GCG GCC GCC GCG CTC CCA 55

Met Ala Ala Ala Leu Pro

-15 -10

GCA TGG CTG TOT CTG CAG TCG AGG GCA AGG ACT CTG CGT GCA TTC TCC

Ala Trp Leu Ser Leu Gln Ser Arg Ala Arg Thr Leu Arg Ala Phe Ser

					20	U				
	GCC Ala								151	
	ACA Thr 25								199	
	ACA Thr								247	
	GCA Ala						 	 	 295	
	TAT Tyr								343	
-	TTT Phe				 				373	

## (2) INFORMATION FOR SEQ ID NO: 218:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 333 base pairs

(3) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 32..331

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99 region 1..300

id R13004

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 114..274

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 54..214 id T80337

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 272..331

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 213..272 'id T80337

est

## (ix) FEATURE:

- (A) NAME/KEY: other (B) LOCATION: 66..106
- (C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 6..46 id T80337 est

## (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 101..278

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 70..247 id T08840 est

## (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 33..113

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 1..81 id T08840 est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 101..249

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 72..220 id HSCOCF041

est

# (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 31..112

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 1..82 id HSCOCF041

est

# (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 247..321

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 5.8

seq LWISACAMLLCHG/SL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 218:

AAGCTAGGAC ATTCTTCTCC TCCTGGCCCT GGACATCAGA ACCCCAGGCT CTCCAGCCTT 60

TGGACTTCAG GACTGACACA AGCAACCTGC TGGGTTCTTA GGCCTTTGGC TTGTACTGAG 120

ACTTACACCA TCAGCTTCCC TGGTCCTGAG ACTTTTGGAC TTGGATTGAG CCACGCTACT 180

GGCATCCCAG GATCTCCAGC TTGCAGACAG CCTGTCGTGG GACTTCACAG CCTCCATAAT 240

TATAGA ATG GCA ATG GTC TCT GCG ATG TCC TGG GTC CTG TAT TTG TGG 288

Met Ala Met Val Ser Ala Met Ser Trp Val Leu Tyr Leu Trp -25 -20 -15

ATA AGT GCT TGT GCA ATG CTA CTC TGC CAT GGA TCC CTT CAG CGG 333

Ile Ser Ala Cys Ala Met Leu Leu Cys His Gly Ser Leu Gln Arg

### (2) INFORMATION FOR SEQ ID NO: 219:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 284 base pairs(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement(2..282)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100 region 59..339

id H10776 est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (64..282)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 73..291 id N94455

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (2..85)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 271..354

id N94455 est

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (107..282)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 58..233 id H64097 est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement(2..120)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 219..337

id H64097

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (38..282)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 60..304 id R98226

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (161..282)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 33..154 id W60134

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement(9..120)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 195..306

id W60134

est

#### (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 51..257

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 5.7

seq LCRLLCLVRLFCC/SS

56

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 219:

ATCAACCATC CAGCTCCCAG CTGGCTAAAC TTTGCCTCCA GTGGTCAAAG ATG GGA Met Gly

AAA GAG TGG GGT TGG CAG GAG ATG GAA AAC GGA GGT GCC CCCA GCA
Lys Glu Trp Gly Trp Gln Glu Met Glu Asn Gly Gly Ala Ala Pro Ala

-65 **-6**0 **-55** 

TGG GGG GCA GGT CCC CCA GTC CAC CCT GCC CCT CCC CCT GTG GAG AAG 15:

Trp Gly Ala Gly Pro Pro Val His Pro Ala Pro Pro Pro Val Glu Lys
-50 -45 -40

ACG CTT AGT TGG GGG TGT GGG TTT GGG CTC CAT TCT GGA TTC GGC GGT

Thr Leu Ser Trp Gly Cys Gly Phe Gly Leu His Ser Gly Phe Gly Gly

-35

-20

TCC GGG GGA GGG GTG GGT CTG TGC CGA TTA CTC TGT CTT GTA CGT TTG

Ser Gly Gly Val Gly Leu Cys Arg Leu Leu Cys Leu Val Arg Leu

-15

-10

-5

TTC TGC TGC TCT TCA ATA TTG TAT CAA CGC CAG GGG

Phe Cys Cys Ser Ser Ile Leu Tyr Gln Arg Gln Gly

# (2) INFORMATION FOR SEQ ID NO: 220:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 361 base pairs(B) TYPE: NUCLEIC ACID(C) STRANDEDNESS: DOUBLE(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 137..358

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97 region 151..372 id N33828

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 2..124

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 14..136 id N33828

est

(ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 138..358

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 147..367

id N34173

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 1..148

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 11..158 id N34173

est

(ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 35..358

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 1..324 id T89546

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 138..337

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 107..306

id H67305

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 42..148

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 12..118

id H67305

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 138..302

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 112..276

id T79378

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 33..145

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 8..120

id T79378

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 317..348

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 293..324

id T79378

est

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide
(B) LOCATION: 167..229

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 5.6

seq LVLSLQFLLLSYD/LF

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 220:

AATGACAACC GACGTTGGAG TTTGGAGGTG CTTGCCTTAG AGCAAGGGAA ACAGCTCTCA TTCAAAGGAA CTAGAAGCCT CTCCCTCAGT GGTAGGGAGA CAGCCAGGAG CGGTTTTCTG 120 GGAACTGTGG GATGTGCCCT TGGGGGCCCG AGAAAACAGA AGGAAG ATG CTC CAG 175 Met Leu Gln -20 ACC AGT AAC TAC AGC CTG GTG CTC TCT CTG CAG TTC CTG CTG CTG TCC 223 Thr Ser Asn Tyr Ser Leu Val Leu Ser Leu Gln Phe Leu Leu Ser -15 -10 TAT GAC CTC TTT GTC AAT TCC TTC TCA GAA CTG CTC CAA AAG ACT CCT 271 Tyr Asp Leu Phe Val Asn Ser Phe Ser Glu Leu Leu Gln Lys Thr Pro GTC ATC CAG CTT GTG CTC TTC ATC ATC CAG GAT ATT GCA GTC CTC TTC 319 Val Ile Gln Leu Val Leu Phe Ile Ile Gln Asp Ile Ala Val Leu Phe 20 AAC ATC ATC ATT TTC CTC ATG TTC TTC AAC ACC TCC CGG 361 Asn Ile Ile Ile Phe Leu Met Phe Phe Asn Thr Ser Arg 35 40

# (2) INFORMATION FOR SEQ ID NO: 221:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 252 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: complement(100..250)
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 95

region 256..406 id W72958

10 W/2938

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 115..250

(C) IDENTIFICATION METHOD: blastn
(D) OTHER INFORMATION: identity 99
region 2..137
id W78821

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 120..250

(C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 99

region 1..131 id AA083784 est

(ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 115..250

(C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 98

region 4..139 id W24219

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 145..250

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97 region 39..144

id C15963

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 114..153

(C) IDENTIFICATION METHOD: blastn
(D) OTHER INFORMATION: identity 95
region 9..48
id C15963

id C1596

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide
(B) LOCATION: 172..243

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 5.5

seq MGVCLLIPGLATA/CI

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 221:

AGCGCGASGY CCGKCTCTCT TGTGCCCTAG CAGATTCCGT CGCTTCTTCC GGAGCCGTAC 60

GT3GTACCGC CCCGCTCGCG GGCGGCCGCG RGGCTTGCTG GGAAGAGAGG CGAACCAGGT 120

CACCTTTCAA GGACCCAGAA GTAGGGTTTT GGCCTAGGTA ACGGGGCAGA G ATG TGG 177
Met Trp

TTC GAG ATT CTC CCC GGA CTC TCC GTC ATG GGC GTG TGC TTG TTG ATT 225
Ph4 Glu Ile Leu Pro Gly Leu Ser Val Met Gly Val Cys Leu Leu Ile

WO 99/06552

216

-20

-15

-10

CCA GGA CTG GCT ACT GCG TGC ATC CGG
Pro Gly Leu Ala Thr Ala Cys Ile Arg
-5

1

(2) INFORMATION FOR SEQ ID NO: 222:

(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 167 base pairs
(B) TYPE: NUCLEIC ACID
(C) STRANDEDNESS: DOUBLE
(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(F) TISSUE TYPE: Brain

(vi) ORIGINAL SOURCE:

(ix) FEATURE:
 (A) NAME/KEY: other

(B) LOCATION: complement (2..103)

(A) ORGANISM: Homo Sapiens

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98 region 48..149 id AA126155

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (98..143)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95 region 7..52 id AA126155

est

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 30..95

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 5.5

seq LADPLXLFPFSEG/LP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 222:

ACTGCTCSTG GAGCTCTGCG CTGGTCTTC ATG CGC CCT AGC CCT CTT TCG GGG

Met Arg Pro Ser Pro Leu Ser Gly

-20 -15

ATA CTG GCC GAC CCC CTC TKC CTT TTC CCC TTT AGT GAA GGC CTC CCC 101

Ile Leu Ala Asp Pro Leu Xaa Leu Phe Pro Phe Ser Glu Gly Leu Pro
-10 -5 1

CGT CGC CGC GCG GCT TCC CGG AGC CGA CTG CAG ACT CCC TCA GCC CGG
Arg Arg Arg Ala Ala Ser Arg Ser Arg Leu Gln Thr Pro Ser Ala Arg

10
15

167

TGT TCC CCG CGT CCG GGG Cys Ser Pro Arg Pro Gly 20

# (2) INFORMATION FOR SEQ ID NO: 223:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 350 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 40..352
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 96

region 30..342

id H15315

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 12..46
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 91

region 1..35

id H15315

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 77..300
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 96

region 1..224

id HUM427H08B

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 22..134
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 91

region 3..115

id AA071651

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 138..326
  - (C) IDENTIFICATION METHOD: blastn

218 . (D) OTHER INFORMATION: identity 95 region 32..220 id R35596 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 65..111 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 91 region 1..47 id W55530 est (ix) FEATURE: (A) NAME/KEY: sig\_peptide (B) LOCATION: 261..341 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 5.4 seg SLMMAQXFIPAVA/KV (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 223: AGGAGGGCTG GACAGCAGCT CAGCTCGCTA GCTGCGCGCT TCCCGGCACA GGCAGTGCCA 60 CTGCGCAGGT TGATCAGCGA AACAGCATCC ATTTTAATCT GCGGGGAGNN CCTGCCTTAC 120 CAGGGCGTTC TCTCCGCCCG CCGGTGGATG CTCCGCGCCT GCSCTCCGCA GCCTCGCTCA 180 GCAGTCCTGC GTTGGGGTCT GCGCCCTAGG ATGCACTGAG ATGGTACATC AGGATAACTG 240 CTCGTATCAG GCACAGAAAA ATG AGA GAG AGT CTA TCA DKS AGA AGT TGG CAC 293 Met Arg Glu Ser Leu Ser Xaa Arg Ser Trp His -25 -20

TTG CCA GCT TCT TTG ATG GCC CAG GKA TTT ATA CCA GCT GTA GCA 341 Leu Pro Ala Ser Leu Met Met Ala Gln Xaa Phe Ile Pro Ala Val Ala -15

AAA GTA GGA 350 Lys Val Gly 1

# (2) INFORMATION FOR SEQ ID NO: 224:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 430 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:

(B) LOCATION: 226..295
(C) IDENTIFICATION METHOD: blastn
(D) OTHER INFORMATION: identity 95
region 121..190
id W07343

#### (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 251..424

(A) NAME/KEY: other

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 5.4

seq LSLHLLATRACYG/IL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 224:

GTTTGGGAGC GAGCAGTTTC CTGCCCAGGG ATGGGGGTCC TGGCTGCACT TCACGGGGGC 120 GGCCCTTTCG TTTCGCTCTG CGTGACAGGT CTCGCTTGAT TGGGTTTCTC ATGGGTSKCT 180 ATTCTTCAAA ATG TCA GGT GTG GTA CCC ACA GCC CCT GAA CAG CCT GCA 289 Met Ser Gly Val Val Pro Thr Ala Pro Glu Gln Pro Ala NGT GAA ATG GAA AAT CAA ACA AAA CCA CCA GAT CCA AGG CCT GAT GCT 337 Xaa Glu Met Glu Asn Gln Thr Lys Pro Pro Asp Pro Arg Pro Asp Ala -45 -40 -35 CCT CCT GAA TAC AGT TCT CAT DBG TTT ACC AGG ACC CCC TGG AAA CAG 385 Pro Pro Glu Tyr Ser Ser His Xaa Phe Thr Arg Thr Pro Trp Lys Gln -25 CTG TCC CTC CAC CTA CTG GCT ACC AGA GCT TGC TAT GGG ATA CTA 430 Leu Ser Leu His Leu Leu Ala Thr Arg Ala Cys Tyr Gly Ile Leu

#### (2) INFORMATION FOR SEQ ID NO: 225:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 387 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: complement (75..325)

(C) IDENTIFICATION METHOD: blastn
(D) OTHER INFORMATION: identity 100 region 82..332 id AA004751 est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement(88..255)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 97

region 153..320 id N27443

est

# (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement(18..105)
- (C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 304..391 id N27443

est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement (258..325)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 100

region 81..148

id N27443

est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement(22..325)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 99

region 80..383

id AA015608

est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement (78..253)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 99

region 165..340

id H09727

est

### (ix) FEATURE:

- (A) NAME/KEY: other
- (3) LOCATION: complement (253..285)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 96

region 132..164

id H09727

est

# (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: complement(49..276)
(C) IDENTIFICATION METHOD: blastn
(D) OTHER INFORMATION: identity 97

region 133..360 id AA027099

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement(269..325)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 83..139 id AA027099

est

# (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 139..369

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 5.3

seq TWVFTCLVFFCFG/LS

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 225:

AGAAACAGGG AGAAG	GAGGAA GGCTAGAAGC	CTGAGCAAGT	GAGGGTAGAA CCTTT	TGGGA 60
CTGGCCTTTG AAGCT	CTGGC CAGGGATGG	GTGGGGGCCA	AAAGGACAGA GCCTG	GTATG 120
TCTTCATAGT CATTO	GAGA ATG TGG AGA Met Trp Arg -75		GGG TGG GGG GTG Gly Trp Gly Val -70	
	AGG GAG ATC CCC Arg Glu Ile Pro -60			
	CCC CCT CTC CCT Pro Pro Leu Pro -45			
	ACC AGG GTC TGG Thr Arg Val Trp -30			
	AGA ACC TGG GTT Arg Thr Trp Val	<del>-</del> -	<del>-</del>	
	MCC TCG CTA GGG Xaa Ser Leu Glÿ 5			387

# (2) INFORMATION FOR SEQ ID NO: 226:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 300 base pairs

(B) TYPE: NUCLEIC ACID
(C) STRANDEDNESS: DOUBLE
(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 123..295

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 121..293 id N78275

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 43..128

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 40..125 id N78275 est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 19..295

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 4..280 id R35388

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 40..295

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 14..269 id W03418

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 29..283

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 27..281 id HSC29H041

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 49..266

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96 region 78..295

id R60376 est

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 184..270

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 5.2

seq FFMLLGSLLPVKI/IE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 226:

AAGACTGCGC GGCCGTWGGG CGTGCAGCGG CGCCAGTCGG CGGACGAGGG GCCCCCGGGA 60 GTTGCTGGAC TGAGACATGA GCCTCCAACT GTGTGGTTGG GCTCGGTAGC ACATCGTGGG 120 ACTTGGGTGT GCGCCCACAG ATGGTTTGGC CCTGCAGTGA CCAGAGCAGC CCAAGCCGCC 180 ACC ATG GTG AAA TTG CTA GTG GCC AAA ATC CTG TGC ATG GTG GGC GTG 228 Met Val Lys Leu Leu Val Ala Lys Ile Leu Cys Met Val Gly Val -25 TTC TTC TTC ATG CTG CTC GGC TCC CTG CTC CCC GTG AAG ATC ATC GAG Phe Phe Phe Met Leu Leu Gly Ser Leu Leu Pro Val Lys Ile Ile Glu -10 -5 300 ACA GAT TTT GAG AAG GCC CCA GGG Thr Asp Phe Glu Lys Ala Pro Gly 5

# (2) INFORMATION FOR SEQ ID NO: 227:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 76 base pairs(B) TYPE: NUCLEIC ACID(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens
(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement(2..73)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 40..111 id HSC39G092

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (2..73)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

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region 37..108 id T89094 est

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

- (B) LOCATION: 11..61
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 5.2

seq IMCLIGLKANASS/ET

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 227:

ATCCTTTTGC ATG CCT GTT TCT ATC ATG TGC TTG ATA GGC CTC AAA GCT

Met Pro Val Ser Ile Met Cys Leu Ile Gly Leu Lys Ala

-15

-5

AAT GCT TCC AGT GAA ACA CAC TCA GGG Asn Ala Ser Ser Glu Thr His Ser Gly 1 5 76

- (2) INFORMATION FOR SEQ ID NO: 228:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 125 base pairs
    - (B) TYPE: NUCLEIC ACID
    - (C) STRANDEDNESS: DOUBLE
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: CDNA
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: other
    - (B) LOCATION: 11..120
    - (C) IDENTIFICATION METHOD: blastn
    - (D) OTHER INFORMATION: identity 100 region 1..110 id HSC3IG111

est

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 48..98
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5.2

seq LLYLVLEKLVSRA/FQ

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 228:

AGATACTAAT CCTTTAAAAA AGTGTAAATG GAGAAAAGTT ATATTTT ATG AAG GTT
Met Lys Val

56

ATT TTG TTG TAT TTA GTA TTG GAA AAG TTG GTT TCC AGA GCA TTT CAG

104

11e Leu Leu Tyr Leu Val Leu Glu Lys Leu Val Ser Arg Ala Phe Gln

-10 -5 1

AAT GTC GAA GCA CCA CAC GGG Asn Val Glu Ala Pro His Gly 125

# (2) INFORMATION FOR SEQ ID NO: 229:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 170 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 81..170
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 97

region 54..143 id T09307

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 29..81
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 92

region 1..53

id T09307

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 12..77
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 98

region 1..66

id AA159859

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 28..75
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 95

region 1..48

id H13321

est

(ix) FEATURE:

. (A) NAME/KEY: other (B) LOCATION: 15..75 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 93 region 10..70 id W02365 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 33..77 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 95 region 1..45 id AA113927 est (ix) FEATURE: (A) NAME/KEY: sig\_peptide (B) LOCATION: 33..89 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 5.1 seg LLLGGRVCXPSLA/VG (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 229: AAGCCGAYYG CTGAAGGCTG GTTTGCGTCG AC ATG GCG GTT ACC CTG AGT CTC Met Ala Val Thr Leu Ser Leu TTG CTG GGC GGG CGC GTT TGC SCG CCG TCA CTC GCT GTG GGT TCG CGA 101 Leu Leu Gly Gly Arg Val Cys Xaa Pro Ser Leu Ala Val Gly Ser Arg CCC GGG GGG TGG CGG GCC CAG GCC CTA TTG GCC GGG AGC CGG ACC CCG 149 Pro Gly Gly Trp Arg Ala Gln Ala Leu Leu Ala Gly Ser Arg Thr Pro 10 15 20 ATT CCG ACT GGG AAC CGG AGG 170 Ile Pro Thr Gly Asn Arg Arg 25 (2) INFORMATION FOR SEQ ID NO: 230: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 263 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR (ii) MOLECULE TYPE: CDNA

(ix) FEATURE:

(vi) ORIGINAL SOURCE:

(A) NAME/KEY: other

(A) ORGANISM: Homo Sapiens
(F) TISSUE TYPE: Brain

(B) LOCATION: 57..261 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 97 region 40..244 id R59037 est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement(184..237)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 100

region 38..91 id R67654 est

#### (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 117..185
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 5.1

seq LLPELGVVTPAQG/PR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 230:

AAGACCATCA ACTATGGAAA GGAGATCTAG GGAACACCGT CTTGAACCCG CCAGGGTTTT

GAGTCTCGGA CCCAGGAGAT CCAACCCTGA CCACCCTCCC AGGATGCAGC AGGGGG ATG 119

TTA AAT CAG ACT TCA GGA AGA ACT TCC TTG CTG CCT GAG TTA GGT GTC 167 Leu Asn Gln Thr Ser Gly Arg Thr Ser Leu Leu Pro Glu Leu Gly Val

-20 -15

GTC ACG CCT GCC CAG GGG CCA AGG AGG CGG GTT TGG TGC GGC CAC TCC 215 Val Thr Pro Ala Gln Gly Pro Arg Arg Arg Val Trp Cys Gly His Ser

AAG GCC AAA GCG AGA AAA TCT TAC TGC GCA CGC GCA ATA GAC TGC CAG Lys Ala Lys Ala Arg Lys Ser Tyr Cys Ala Arg Ala Ile Asp Cys Gln 15 20

# (2) INFORMATION FOR SEQ ID NO: 231:

# (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 430 base pairs
- (B) TYPE: NUCLEIC ACID
- (C) STRANDEDNESS: DOUBLE
- (D) TOPOLOGY: LINEAR

#### (ii) MOLECULE TYPE: CDNA

- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other

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(B) LOCATION: 99..416 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 99 region 1..318 id T31969 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 49..334 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 99 region 2..287 id HSB03B072 (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: complement (2..57) (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 100 region 1..56 id W51830 est (ix) FEATURE: (A) NAME/KEY: sig\_peptide (B) LOCATION: 26..262 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 5 seq SFLGFSAPTPIQA/LT (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 231: ATGTGATGAT CCGGAGGCTG GGGAG ATG ACA TCA GAA AAC CTG GTC CAA ACT 52 Met Thr Ser Glu Asn Leu Val Gln Thr GCT CCA AAA AAG AAG AAA AAT AAA GGG AAA AAA GGG TTG GAG CCT TCT 100 Ala Pro Lys Lys Lys Asn Lys Gly Lys Lys Gly Leu Glu Pro Ser -65-60 CAG AGC ACT GCT GCC AAG GTG CCC AAA AAA GCG AAG ACA TGG ATT CCT 148 Gln Ser Thr Ala Ala Lys Val Pro Lys Lys Ala Lys Thr Trp Ile Pro GAA GTT CAT GAT CAG AAA GCA GAT GTG TCA GCT TGG AAG GAC CTG TTT 196 Glu Val His Asp Gln Lys Ala Asp Val Ser Ala Trp Lys Asp Leu Phe GTT CCC AGG CCG GTT CTC CGA GCA CTC AGC TTT CTA GGC TTC TCT GCA 244 Val Pro Arg Pro Val Leu Arg Ala Leu Ser Phe Leu Gly Phe Ser Ala -15 CCC ACA CCA ATC CAA GCC CTG ACC TTG GCA CCT GCC ATC CGT GAC AAA 292 Pro Thr Pro Ile Gln Ala Leu Thr Leu Ala Pro Ala Ile Arg Asp Lys

CTG GAC ATC CTT GGG GCT GCT GAG ACA GGA AGT GGG AAA ACT CTT GCC

Leu Asp Ile Leu Gly Ala Ala Glu Thr Gly Ser Gly Lys Thr Leu Ala

340

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TTT GCC ATC CCA ATG ATT CAT GCG GTG TTG CAG TGG CAG AAG AGG AAT

TTT GCC ATC CCA ATG ATT CAT GCG GTG TTG CAG TGG CAG AAG AGG AAT

Phe Ala Ile Pro Met Ile His Ala Val Leu Gln Trp Gln Lys Arg Asn

30 35 40

GCT GCC CCT CCA AGT AAC ACC GAA GCA CCA CCT GGA GAG
Ala Ala Pro Pro Pro Ser Asn Thr Glu Ala Pro Pro Gly Glu
45 50 55

#### (2) INFORMATION FOR SEQ ID NO: 232:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 252 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 1..37
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 97 region 9..45 id W84513 est
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 16..84
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.9 seq WHXLIPLTWACMA/RQ
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 232:
- AATCTTCTCC GCGCT ATG GCT GCG TTC GGC CGT CAG SCW TTS ART TGG CAC

  Met Ala Ala Phe Gly Arg Gln Xaa Xaa Xaa Trp His

  -20

  -15
- CKY CTG ATC CCC CTC ACC TGG GCC TGT ATG GCT AGG CAG ACT CCT CAT

  Xaa Leu Ile Pro Leu Thr Trp Ala Cys Met Ala Arg Gln Thr Pro His

  -10 -5 1 5
- CTT GGA GAA CAG AGA AGG ACG ACA GCT TCT TTG TKG CGC AAA CTG ACT
  Leu Gly Glu Gln Arg Arg Thr Thr Ala Ser Leu Xaa Arg Lys Leu Thr
  10 15 20
- ACA GCC TCC AAT GGA GGG GTC ATT GAG GAG TTA TCT TGT GTK AGA TCC

  Thr Ala Ser Asn Gly Gly Val Ile Glu Glu Leu Ser Cys Val Arg Ser

  25

  30

  35

AAT AAC TAT GTG CAG GAA CCA GAG TGC AGG AGG AAT CTT GTT CAG TGC 243 Asn Asn Tyr Val Gln Glu Pro Glu Cys Arg Arg Asn Leu Val Gln Cys 45

CTC CTC TGG Leu Leu Trp 55

252

#### (2) INFORMATION FOR SEQ ID NO: 233:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 347 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 44..187
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 96

region 1..144 id AA151232

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 187..285
  - (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 97
    - region 143..241

id AA151232

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 314..349
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 100

region 272..307

id AA151232

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (3) LOCATION: 39..225
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 95

region 2..138

id AA040887

est

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide (B) LOCATION: 144..314 (C) IDENTIFICATION METHOD: Von Heijne matrix (D) OTHER INFORMATION: score 4.9 seq GWFLSGCPHGSSA/TW (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 233: AATGGGGATG TTGAATTTGG AAATTGGAGG GGACGCTGGT GGWYKKATTG GGTGCAAGGA 60 GTTGGTGTTG ATGGAGGAGC AGGASRCCAG AGTCCCAGCC CTGGAACCGT TCAGAGTGGA GCAGGCACCA CCTGTAATCT ACT ATG TCC CTG ACT TCA TCT CCA AAG AAG AGG Met Ser Leu Thr Ser Ser Pro Lys Lys Arg -55 -50 AGG AGT ATT TGC TTC GAC AGG TTT TTA ATG CCC CAA AGC CAA AGT GGA 221 Arg Ser Ile Cys Phe Asp Arg Phe Leu Met Pro Gln Ser Gln Ser Gly -45 -40 CCC AGC TCT CTG GGA GAA AGT TAC AGA ACT GGG GTG GGC TTC CTC ATC Pro Ser Ser Leu Gly Glu Ser Tyr Arg Thr Gly Val Gly Phe Leu Ile -25 CCC GAG GGA TGG TTC CTG AGC GGC TGC CCC CAT GGC TCC AGC GCT ACG 317 Pro Glu Gly Trp Phe Leu Ser Gly Cys Pro His Gly Ser Ser Ala Thr -10 TGG ACA AAG TGT CAA ACC TCA GCC TCT TTG 347 Trp Thr Lys Cys Gln Thr Ser Ala Ser Leu 5 10 (2) INFORMATION FOR SEQ ID NO: 234: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 227 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR (ii) MOLECULE TYPE: CDNA (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: 115..226 (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 95 region 135..246 id HSC0GF021 est

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide (B) LOCATION: 90..206

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.8 seg SLXFCLSPPPSPS/LR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 234:

AAACCCCATA CCCCCTCCCC ATCTTGTGAT CACCCTCATT ACCTCTTCTG GGCCCCCTGT

GGACCTGCGT TGACCCAGCA TGGGCTACA ATG GGG GAG TTG GGT AAT CGC TCC Met Gly Glu Leu Gly Asn Arg Ser

-35

CGT TGC ATC CTG TTT CTG TCT GAA AAC CCT TGT CTT TCT GAA TCC ATC 161 Arg Cys Ile Leu Phe Leu Ser Glu Asn Pro Cys Leu Ser Glu Ser Ile -30 -25

209

TTT CAG TCT CTS RCA TTC TGT CTT TCC CCT CCT CCT TCA CCT TCC CTC Phe Gln Ser Leu Xaa Phe Cys Leu Ser Pro Pro Pro Ser Pro Ser Leu -10 -5

CGT CCC TCT CCC TCA CGG Arg Pro Ser Pro Ser Arg

227

5

(2) INFORMATION FOR SEQ ID NO: 235:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 430 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 101..355

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 83..337 id AA057242

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 57..101

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100 region 40..84

id AA057242

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 357..400

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 338..381 id AA057242

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 18..51

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 1..34 id AA057242 est

# (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 400..431

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 382..413 id AA057242

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 84..218

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 73..207 id R09808

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 10..51

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 1..42 id R09808

est

#### (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 98..376

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.7

seq VLLLRQXFAQAEK/WY

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 235:

AATTTTCYGT GGTCCAACTA CCCTCGGCGA TCCCAGGCTT GGCGGGGCAC CGCCTGGCCT 60

CTCCCGTTCC TTTAGGCTGC CGCCGCTGCC TGCCGCC ATG GCA GAG TTG GGC CTA 115 Met Ala Glu Leu Gly Leu

-90

AAT GAG CAC CAT CAA AAT GAA GTT ATT AAT TAT ATG CGT TTT GCT CGT Asn Glu His His Gln Asn Glu Val Ile Asn Tyr Met Arg Phe Ala Arg

-85 -80 -75

TCA AAG AGA GGC TTG AGA CTC AAA ACT GTA GAT TCC TGC TTC CAA GAC 211 Ser Lys Arg Gly Leu Arg Leu Lys Thr Val Asp Ser Cys Phe Gln Asp CTC AAG GAG AGC AGG CTG GTG GAG GAC ACC TTC ACC ATA GAT GAA GTC 259 Leu Lys Glu Ser Arg Leu Val Glu Asp Thr Phe Thr Ile Asp Glu Val -50 -45 TCT GAA GTC CTC AAT GGA TTA CAA GCT GTG GTT CAT AGT GAG GTG GAA 307 Ser Glu Val Leu Asn Gly Leu Gln Ala Val Val His Ser Glu Val Glu -35 -30 TCT GAG CTC ATC AAC ACT GCC TAT ACC AAT GTG TTA CTT CTG CGA CAG 355 Ser Glu Leu Ile Asn Thr Ala Tyr Thr Asn Val Leu Leu Leu Arg Gln NTG TTT GCA CAA GCT GAG AAG TGG TAT CTT AAG CTA CAG ACA GAC ATC 403 Xaa Phe Ala Gln Ala Glu Lys Trp Tyr Leu Lys Leu Gln Thr Asp Ile TCT GAA CTT GAA AAC CGA GAA TTA TTA 430 Ser Glu Leu Glu Asn Arg Glu Leu Leu 15

# (2) INFORMATION FOR SEQ ID NO: 236:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 344 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 20..231
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 99

region 1..212 id N33729

est

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 135..281
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.6

seq SWAVGLLYAVAQG/SK

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 236:

AATTAGO	GAG (	GCCA'	rGGG(	GG AA	AAAA	STCT	A ACT	rggco	GGAA	CTC	CTGG	GAA (	CTGG	GCGAT	60
GGGCTCT	TAG	TATCO	GAG	SA TI	rggac	CCAT	г сто	SATT	ATT	CCT	GAAA!	TTC (	CTTAC	STCTCT	120
CCTGTGT	TGG (	GGAA			ACC Thr										170
TGT CCA Cys Pro															218
GAA GAC Glu Asp -20	Ala														266
GCA GTG Ala Val -5															314
CTT NGT Leu Xaa															344

# (2) INFORMATION FOR SEQ ID NO: 237:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 419 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 116..419
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 98

region 122..425

id W68799

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 18..117
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 97

region 1..100

id W68799

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 18..209

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 93

region 1..192 id W49697 est

# (ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 199..290

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 183..274

id W49697

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 291..367

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 276..352

id W49697 est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 387..417

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 374..404

id W49697

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 48..419

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 1..372

id AA149518

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 171..414

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 116..359

id W17032

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 57..174

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 1..118

id W17032

est

# (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 78..386
(C) IDENTIFICATION METHOD: blastn
(D) OTHER INFORMATION: identity 98
region 1..309
id W78749
est

(ix) FEATURE:
(A) NAME/KEY: other
(B) LOCATION: 386..419
(C) IDENTIFICATION METHOD: blastn
(D) OTHER INFORMATION: identity 100
region 310..343
id W78749
est

#### (ix) FEATURE:

- (A) NAME/KEY: sig peptide
- (B) LOCATION: 180..383
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 4.6

seq LPFSLVSMLVTQG/LV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 237:

AAGACAGGTG GGGTACTCGG GAAGCTGGAG CGGGCCGGCG GTGCAGTCAC GGGGGAGCGA 60 GGCCTGCTGG GCTTGGCAAC GAGGGACTCG GCCTCGGAGG CGACCCAGAC CACACAGACA 120 CTGGGTCAAG GAGTAAGCAG AGGATAAACA ACTGGAAGGA GAGCAAGCAC AAAGTCATC 179 ATG GCT TCA GCG TCT GCT CGT GGA AAC CAA GAT AAA GAT GCC CAT TTT 227 Met Ala Ser Ala Ser Ala Arg Gly Asn Gln Asp Lys Asp Ala His Phe -60 CCA CCA CCA AGC AAG CAG AGC CTG TTG TTT TGT CCA AAA TMH NNA CTG 275 Pro Pro Pro Ser Lys Gln Ser Leu Leu Phe Cys Pro Lys Xaa Xaa Leu -45 CAC ATC CAC AGA GCA GAG ATC TCA AAG ATT ATG CGA GAA TGT CAG GAA His Ile His Arg Ala Glu Ile Ser Lys Ile Met Arg Glu Cys Gln Glu -30 GAA AGT TTC TGG AAG AGA GCT CTG CCT TTT TCT CTT GTA AGC ATG CTT 371 Glu Ser Phe Trp Lys Arg Ala Leu Pro Phe Ser Leu Val Ser Met Leu GTC ACC CAG GGA CTA GTC TAC CAA GGT TAT TTG GCA GCT AAT TCT AGA 419 Val Thr Gln Gly Leu Val Tyr Gln Gly Tyr Leu Ala Ala Asn Ser Arg

# (2) INFORMATION FOR SEQ ID NO: 238:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 274 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

#### (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens(F) TISSUE TYPE: Brain

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement (37..269)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 99

region 2..234 id AA147071 est

# (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement(2..31)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 93

region 239..268 id AA147071 est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement (37..269)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 99

region 58..290

id H98153

est

# (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement(2..31)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 93

region 295..324

id H98153

est

# (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement (37..269)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 98

region 59..291

id N49401

est

# (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement(2..31)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 93

region 296..325

id N49401

est

274

iх				

(A) NAME/KEY: other

(B) LOCATION: complement (87..269)

(C) I-DENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 37..219

id N80022

est

# (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 62..268

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.5

seq FILSLCVLCIVLT/TG

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 238:

#### AATTAAGTCA KDATACAAAT CAGCACAGAT AACGDMAATG TTTCCAATAT WWTAAAATGT

A ATG TTA CTT ATG AAA AGT ATT TTG CTT AAG GTT GTG TGT GTA TTG TGT 109 Met Leu Leu Met Lys Ser Ile Leu Leu Lys Val Val Cys Val Leu Cys -65 -60

ATA TAC CTC AAG TTC AAG TTA ATG GCA TTG ATT TAT GTT CCA GAC AAA 157 Ile Tyr Leu Lys Phe Lys Leu Met Ala Leu Ile Tyr Val Pro Asp Lys

AAT AAC ACA AAT AAT AAT ATC CTT CGT TAT AAC CAC AAT GAG ATA AGT 205 Asn Asn Thr Asn Asn Asn Ile Leu Arg Tyr Asn His Asn Glu Ile Ser -35 -30

ATT GGC ATT AGT GTT CAG TGC CAT TTT ATA CTT TCT CTC TGT GTT CTC 253 Ile Gly Ile Ser Val Gln Cys His Phe Ile Leu Ser Leu Cys Val Leu -20 -15

TGT ATT GTA CTA ACC ACT GGG Cys Ile Val Leu Thr Thr Gly

### (2) INFORMATION FOR SEQ ID NO: 239:

# (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 249 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

### (ii) MOLECULE TYPE: CDNA

# (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 100..249

(C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 94 region 20..169 id N41898 est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 113..249

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 38..174 id H69272

est

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 100..147

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.5

seg RLLLRRFLASVIS/RK

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 239:

AGTTATGTAC GTTCCCCCC CCGAGGAAGT GAYGACAGGC GTGCCCTTGA CAGGCAGGGA 60

GGGCTAGGCT GTGCATCCCT CCGCTCGCAT TGCAGGGAG ATG GCT CAG CGA CTT 114 Met Ala Gln Arg Leu

CTT CTG AGG AGG TTC CTG GCC TCT GTC ATC TCC AGG AAG CCC TCT CAG 162 Leu Leu Arg Arg Phe Leu Ala Ser Val Ile Ser Arg Lys Pro Ser Gln -10

GGT CAG TGG CCA CCC CTC ACT TCC AGA GCC CTG CAG ACC CCA YAA TGC 210 Gly Gln Trp Pro Pro Leu Thr Ser Arg Ala Leu Gln Thr Pro Xaa Cys 10 15

AGT YCT GGT GGC CTG ACT GTA ACA CCC AAC CCA AGC CGG 249 Ser Xaa Gly Gly Leu Thr Val Thr Pro Asn Pro Ser Arg

25 30

(2) INFORMATION FOR SEQ ID NO: 240:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 310 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 51..209

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 49..207 id N56053

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 2..54

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 1..53 id N56053

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 211..246

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 208..243

id N56053 est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 275..307

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 270..302

id N56053

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 51..178

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 44..171

id R59444

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 212..275

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 203..266

id R59444

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 7..54

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 1..48 id R59444

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 274..308

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 266..300 id R59444

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 178..209

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 170..201

id R59444

est

(ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 51..178

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 45..172 id AA156837

est

(ix) FEATURE:

(A) NAME/KEY: other

(3) LOCATION: 178..246

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 171..239 id AA156837

est

(ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 247..308

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 239..300 id AA156837

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 6..54

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 1..49

id AA156837

est

(ix) FEATURE:

(A) NAME/KEY: other

(3) LOCATION: 51..178

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 56..183

id N88392

est

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(ix) FEATURE:
```

(A) NAME/KEY: other (B) EOCATION: 13..54

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 19..60 id N88392 est

# (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 247..285

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 249..287 id N88392

est

#### (ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 211..246

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 214..249 id N88392

est

#### (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 179..209

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 183..213

id N88392

est

#### (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 7..209

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 1..203

id R18752

est

# (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 211..246

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 204..239

id R18752

est

#### (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION:  $2..\overline{2}32$ 

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.4

# seq FEARIALLPLLQA/ET

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 240:

A ATG GCG GC Met Ala Al	la Ser Lys V	TG AAG CAG GA al Lys Gln As -70	AC ATG CCT CC sp Met Pro Pr	R MCG GGG GG o Xaa Gly G: -65	GC TAT 49 ly Tyr
			CCG CGT CGA Pro Arg Arg		
			ACC CTG ATC Thr Leu Ile -35		
			AGG CGC CTA ( Arg Arg Leu ( -20		
			CTG TTA CAG		
			AAC CTG GAG ( Asn Leu Glu ( 15		
ATC ATG AAG Ile Met Lys 20	GAC GTG CCC Asp Val Pro 25				310

# (2) INFORMATION FOR SEQ ID NO: 241:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 388 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 93..257
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 99 region 103..267 id H87397 est
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide (B) LOCATION: 158..319

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.3 seq LLSLAILSHISTP/GC

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 241:

AGACAAAGAG AAGGCAAAAT SAGTTTGTGT CCCTGAGTTG CTAAGTGGAG AAGAAACGTC 60 CACCAACCAG GAAACACCTG CCTCCAACTG TTAATAGGTC TGTGAAATGT GCTTTGTTTC 120 TGGTCAGCAT GGACACCCGC TTTAATAGTG GCTTCAG ATG AGG CAC CTT GTG ACA 175 Met Arg His Leu Val Thr GAG GAG CTC TTC CCC TGC AGC AAC CTT GAA GAT GTT GTG GAA GAC AAT 223 Glu Glu Leu Phe Pro Cys Ser Asn Leu Glu Asp Val Val Glu Asp Asn -45 -40 AGC CAT TCT TAC TTC ACT CTG AGG ATC ACG ATG GCG TGC AAG GGT GTG 271 Ser His Ser Tyr Phe Thr Leu Arg Ile Thr Met Ala Cys Lys Gly Val -25 CCA AGC ACA TTG CTA TCT TTG GCC ATT CTC TCT CAC ATT AGT ACA CCT 319 Pro Ser Thr Leu Leu Ser Leu Ala Ile Leu Ser His Ile Ser Thr Pro -10 GGA TGT GAA TGG CAC GTT ATC TAT GTA AGC AGT BAT GGT CTC TAT CTT 367 Gly Cys Glu Trp His Val Ile Tyr Val Ser Ser Xaa Gly Leu Tyr Leu

GTG GTA GAA ATG ACA GAC CGG Val Val Glu Met Thr Asp Arg 20

10

388

#### (2) INFORMATION FOR SEQ ID NO: 242:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 391 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 108..392
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 98 region 104..388 id T08101

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 32..110

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 1..79 id T08101 est

# (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 108..392

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 39..323 id T27149

est

# (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 113..392

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 30..309 id H06555

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 108..316

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 90..298

id HSC3CC081

est

#### (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 60..110

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 15..65 id HSC3CC081

est

#### (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 105..316

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 58..269

id T74159

est

### (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 76..105

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 93

region 5..34

id T74159

est

WO 99/06552 247

(ix)	) FEATURE:	

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 152..379
- (C) PDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 4.3

seq FRLLXVFAYGTYA/DY

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 242:

AAGTGGCCAG AGCGACTCTT CAGGGAGGTG GCAGGAAAGG CTTGGAACAG CTGCCGGAGG 60 TGACGGAGCG GCGGCCCGC CCGGTGCGCT GGAGGTCGAA GCTTCCAGCT CTGGACATCC 120 TGAGCCCAAG TCCCCCACAC TCAGTGCAGT G ATG AGT GCG GAA GTG AAG GTG Met Ser Ala Glu Val Lys Val -75 ACA GGG CAG AAC CAG GAG CAA TTT CTG CTC CTA GCC AAG TCG GCC AAG 220 Thr Gly Gln Asn Gln Glu Gln Phe Leu Leu Leu Ala Lys Ser Ala Lys GGG GCA GCG CTG GCC ACA CTC ATC CAT CAG GTG CTG GAG GCC CCT GGT 268 Gly Ala Ala Leu Ala Thr Leu Ile His Gln Val Leu Glu Ala Pro Gly GTC TAC GTG TTT GGA GAA CTG CTG GAC ATG CCC AAT GTT AGA GAG CTG 316 Val Tyr Val Phe Gly Glu Leu Leu Asp Met Pro Asn Val Arg Glu Leu -30 GCT GAG AGT NAC TTT GCC TCT ACC TTC CGG CTG CTC AMA GTG TTT GCT 364 Ala Glu Ser Xaa Phe Ala Ser Thr Phe Arg Leu Leu Xaa Val Phe Ala -15 -10 TAT GGG ACA TAC GCT GAC TAC TWA GCT 391

### (2) INFORMATION FOR SEQ ID NO: 243:

Tyr Gly Thr Tyr Ala Asp Tyr Xaa Ala

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 299 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 47..248
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 97 region 15..216

id HUM429E03B est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 244..299

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 211..266 id HUM429E03B

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 133..299

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 107..273 id T80259

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 47..139

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 92

region 22..114 id T80259

est.

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 48..292

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 1..245 id T31768

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 111..299

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 47..235

id N32697

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 64..106

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 1..43

id N32697

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 74..299

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94

region 1..226 id N44613 est

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(A) NAME/KEY: sig\_peptide

(B) LOCATION: 165..266

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.3

seq QLFAFLNLLPVEA/DI

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 243:

ACTTCCGCTT CGCCTAGGTG TTGTCGTCCC TGCTAGTACT CCGGGCTGGG GGTCGGTGCG GATATTCAGT CATGAAATCA SGGTAGGGAC TTCTCCCGCA GCGACGCGGC TGGCAAGACT 120 GTTTGTGTWG CGGGGGCCGG ACTTCAAGGT GATTTTACAA CGAG ATG CTG CTC TCC Met Leu Leu Ser ATA GGG ATG CTC ATG CTG TCA GCC ACA CAA GTS TAS ACC ATC TTG AST 224 Ile Gly Met Leu Met Leu Ser Ala Thr Gln Val Xaa Thr Ile Leu Xaa -20 GTC CAG CTC TTT GCA TTC TTA AAC CTA CTG CCT GTA GAA GCA GAC ATT 272 Val Gln Leu Phe Ala Phe Leu Asn Leu Leu Pro Val Glu Ala Asp Ile -10 KTA GCA TAT AAC TTT GAA AAT GCA TCT 299 Xaa Ala Tyr Asn Phe Glu Asn Ala Ser 5 10

# (2) INFORMATION FOR SEQ ID NO: 244:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 312 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: complement (115..313)
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 98

region 1..199 id H19659

est

- (ix) FEATURE:
  - (A) NAME/KEY: other

250 (B) LOCATION: complement(2..102) (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 97 region 212..312 id H19659 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: complement(115..313) (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 97 region 1..199 id R72881 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: complement(115..290) (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 98 region 1..176 id H50517 (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: complement (44..102) (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 100 region 189..247 id H50517 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: complement(115..302) (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 96 region 36..223 id H41556 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: complement (115..313) (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 95 region 2..200 id R71794 est (ix) FEATURE: (A) NAME/KEY: other

- (B) LOCATION: complement (44..102)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 96 region 213..271 id R71794 · est

251

(ix	) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 229..276

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.2

seq EVVSLSYCGVSWG/RI

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 244:

ACATTTCTGC TCAGATTCCC GCCATCTCCA TTGCATTCAT GTACTACCCT CAGTCTACAC AGGAAAAATA AGTGGGGGCA GGTTTGGAGA GCTGCTTCCA GTGGATAGTT GATGAGAATC 180 CTGACCAAAG GAAGGCACCC TTGACTGTYG GGATAGACAG ATGGACCT ATG GGG TGG 237 Met Gly Trp -15 GAG GTG GTG TCC CTT TCA TAC TGT GGT GTC TCT TGG GGA AGG ATC TCC 285 Glu Val Val Ser Leu Ser Tyr Cys Gly Val Ser Trp Gly Arg Ile Ser -10

CCG AAT CTC AAT AAA CCA GTG AAC AGG Pro Asn Leu Asn Lys Pro Val Asn Arg 5

312

### (2) INFORMATION FOR SEQ ID NO: 245:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 252 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 41..210
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 98 region 77..246 id R59124

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 37..132
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.2 seq CWELFCLEHGIQA/DG
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 245:

AAA(	GCTG <i>I</i>	AGA (	GKG	CGCGC	GG CC	GAGG	ACAGO	C GGC	CASR			ATA Ile	 54
	CAC His -25												102
	TTC Phe												150
	GCT Ala												198
	ACT Thr												246
	AAA Lys 40												252

#### (2) INFORMATION FOR SEQ ID NO: 246:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 172 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 82..168

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 107..193 id AA088577

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 31..71

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 92

region 53..93 id AA088577

est

(ix) FEATURE:

```
(A) NAME/KEY: other
           (B) LOCATION: 31..168
           (C) IDENTIFICATION METHOD: blastn
           (D) OTHER INFORMATION: identity 97
                                    region 24..161
                                    id R16448
     (ix) FEATURE:
           (A) NAME/KEY: other
           (B) LOCATION: 53..168
           (C) IDENTIFICATION METHOD: blastn
           (D) OTHER INFORMATION: identity 99
                                    region 23..138
                                    id AA094092
                                    est
     (ix) FEATURE:
           (A) NAME/KEY: other
           (B) LOCATION: 31..163
           (C) IDENTIFICATION METHOD: blastn
           (D) OTHER INFORMATION: identity 96
                                    region 24..156
                                    id R18030
                                    est
     (ix) FEATURE:
           (A) NAME/KEY: other
           (B) LOCATION: 60..168
           (C) IDENTIFICATION METHOD: blastn
           (D) OTHER INFORMATION: identity 97
                                    region 43..151
                                    id W00599
                                    est
      (ix) FEATURE:
           (A) NAME/KEY: other
           (B) LOCATION: 29..70
           (C) IDENTIFICATION METHOD: blastn
           (D) OTHER INFORMATION: identity 95
                                    region 13..54
                                    id W00599
                                    est
      (ix) FEATURE:
            (A) NAME/KEY: sig_peptide
            (B) LOCATION: 35..109
            (C) IDENTIFICATION METHOD: Von Heijne matrix
            (D) OTHER INFORMATION: score 4.2
                                    seq LDLLRGLPRVSLA/NL
      (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 246:
AAGGGCGCCC TTGAAAGTTC TTGGATCTGC GGGT ATG GCC GGT CCC TTG CAG GGC 55
                                      Met Ala Gly Pro Leu Gln Gly
                                       -25
                                                           -20
GGT GGG GCC CGG GCC CTG GAC CTA CTC CGG GGC CTG CCG CGT GTG AGC
```

Gly Gly Ala Arg Ala Leu Asp Leu Leu Arg Gly Leu Pro Arg Val Ser

-10

-15

CTG GCC AAC TTA AAG CCG AAT CCC GGC TCC AAG AAA CCG GAG AGA AGA
Leu Ala Asn Leu Lys Pro Asn Pro Gly Ser Lys Lys Pro Glu Arg Arg

1 5 10

CCA AGA GGT CGG AGA AGG TGG
Pro Arg Gly Arg Arg Arg Trp
15 20

#### (2) INFORMATION FOR SEQ ID NO: 247:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 359 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 52..360
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 98

region 1..309 id HSC1ED081

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 171..316
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 98

region 146..291 id AA143136

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 31..165
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 97

region 6..140

id AA143136

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 310..341
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 96

region 286..317

id AA143136

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	AAA															359
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(ii) MOLECULE TYPE: CDNA

#### '(vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo Sapiens
- (F) TISSUE TYPE: Brain

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 10..280
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 98

region 17..287 id AA082102

est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 72..224
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 98

region 30..182 id R10417

est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 221..280
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 100

region 180..239

id R10417

est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 47..280
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 99

region 2..235

id W73318

est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 42..224
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 97

region 1..183

id R08733

est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 237..269
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 93

region 198..230

id R08733

WO 99/06552	257	PCT/IB98/01230

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 39..110
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.2

seg SLPALALSLRASP/RX

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 248:

AAGTGCGTGC GCGGCGACTG CGACGGGCAG TGGCAGTC ATG GCG GTT CAG TGC GTG Met Ala Val Gln Cys Val

AGG TTG GCG CGG CGC AGT CTT CCT GCT TTG GCG TTG TCT CTC AGG GCA 104 Arg Leu Ala Arg Arg Ser Leu Pro Ala Leu Ala Leu Ser Leu Arg Ala -15 -10

TCT CCC CGG KTG TTG TGC ACA GCC ACG AAA CAA AAG AAC AGT GGC CAG 152 Ser Pro Arg Xaa Leu Cys Thr Ala Thr Lys Gln Lys Asn Ser Gly Gln 5

AAC CTG GAA GAG GAC ATG GGT CAG AGT GAA CAG AAG GCA GAT CCT CCT 200 Asn Leu Glu Glu Asp Met Gly Gln Ser Glu Gln Lys Ala Asp Pro Pro

GCT ACA GAG AAG ACC CTC CTG GAA GAG AAG GTC AAG TTG GAG GAA CAG 248 Ala Thr Glu Lys Thr Leu Leu Glu Glu Lys Val Lys Leu Glu Glu Gln

CTG AAG GAG ACT GTG GAA AAA TAT AAA CGA GCG AGG 284 Leu Lys Glu Thr Val Glu Lys Tyr Lys Arg Ala Arg 50

#### (2) INFORMATION FOR SEQ ID NO: 249:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 307 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: complement(34..74)
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 92

region 271..311 id T05270

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide

(B) LOCATION: 182..292

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.2

seq RLMHHYLSTPTSA/RP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 249:

AAAGGCTGCC CTGTGGCACC ACAATCTAAG CTCAGGGCAT AAAACCCCTT GTGGCTTTGA 60

TGGAATCCAG GGCTCAGACC ATAAAACCCC TCGTGGCCTT TTGAATGTGC ACCGACTTGC 120

TGGCTCCTTG CTTCTTGCTC TCCCAGAATC GTAAATTGAT TGTATCTTGA GTTGGAAGAA 180

C ATG TTC TCC ATT ATC TCA CGT AGC AGA GCA TGT TCC ATG TAC TTC AAA 229

Met Phe Ser Ile Ile Ser Arg Ser Arg Ala Cys Ser Met Tyr Phe Lys

-35

-30

-25

GAA AAT GCT AAA CCG TCA CAG CTA CGC TTG ATG CAC CAC TAC CTT TCT 277
Glu Asn Ala Lys Pro Ser Gln Leu Arg Leu Met His His Tyr Leu Ser
-20 -15 -10

ACC CCC ACA TCC GCA CGT CCT CAC CTG 307
Thr Pro Thr Ser Ala Arg Pro His His Leu
-5 5 5

#### (2) INFORMATION FOR SEQ ID NO: 250:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 212 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: complement(1..209)
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 99

region 125..333

id H40205

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: complement (80..209)
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 96

region 131..260

id H03462

est

(ix) FEATURE:

```
259
      (A) NAME/KEY: other
      (B) LOCATION: complement (52..90)
      (C) IDENTIFICATION METHOD: blastn
      (D) OTHER INFORMATION: identity 97
                              region 251..289
                              id H03462
                              est
(ix) FEATURE:
      (A) NAME/KEY: other
      (B) LOCATION: complement (17..54)
      (C) IDENTIFICATION METHOD: blastn
      (D) OTHER INFORMATION: identity 97
                              region 288..325
                              id H03462
                              est
(ix) FEATURE:
      (A) NAME/KEY: other
      (B) LOCATION: complement (17..209)
      (C) IDENTIFICATION METHOD: blastn
      (D) OTHER INFORMATION: identity 95
                              region 130..322
                              id R05443
                              est
(ix) FEATURE:
      (A) NAME/KEY: other
      (B) LOCATION: complement(128..209)
      (C) IDENTIFICATION METHOD: blastn
      (D) OTHER INFORMATION: identity 98
                              region 143..224
                              id T52770
                              est
(ix) FEATURE:
      (A) NAME/KEY: other
      (B) LOCATION: complement (80..128)
      (C) IDENTIFICATION METHOD: blastn
      (D) OTHER INFORMATION: identity 100
                              region 225..273
                              id T52770
                              est
(ix) FEATURE:
      (A) NAME/KEY: other
      (B) LOCATION: complement (43..74)
      (C) IDENTIFICATION METHOD: blastn
      (D) OTHER INFORMATION: identity 96
                              region 281..312
                              id T52770
                              est
(ix) FEATURE:
      (A) NAME/KEY: other
      (B) LOCATION: complement (57..209)
      (C) IDENTIFICATION METHOD: blastn
      (D) OTHER INFORMATION: identity 98
```

region 143..295

id AA037595

WO 99/06552	260	РСТ/ІВ98/01236
(ix) FEATURE:		

(A) NAME/KEY: sig\_peptide
(B) LOCATION: 108..155

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.1

seq LLPATSLAGPVLS/TL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 250:

ACTTCTTGTG GACTCACCAA GAGAAACAAA AGGAAGCCTG CACCATTGTA GCCCTGAACT 60

CTTTTCTGGG CACCTGAATC CCAGGAACCC TCAATGAGGT CTTCAAG ATG AAG AGA Met Lys Arg -15

CTG CTG CCA GCT ACC AGC CTG GCT GGC CCT GTC CTG TCC ACC CTC ATT Leu Leu Pro Ala Thr Ser Leu Ala Gly Pro Val Leu Ser Thr Leu Ile -10 -5 1

GCC CCA ACT CCC ATG TTG TTT TGT GAA GAT AAA AGC TGG GAT CCT GGG Ala Pro Thr Pro Met Leu Phe Cys Glu Asp Lys Ser Trp Asp Pro Gly

#### (2) INFORMATION FOR SEQ ID NO: 251:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 357 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 108..308

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 116..316 id HSC2TH021

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 16..99

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 24..107 id HSC2TH021

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 30..92

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 92

region 72..134 id W54529 est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 119..352

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 23..256 id R59681

est

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 64..273

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4

seq IAVLYLHLYDVFG/DP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 251:

AACTGTCCGG GGCTGCGGGG CTTGCTTCCG GCGTCATGGC TC									TCA	AAGG	GCC 1	TCC	CGAATC	60		
CTT						AAT Asn -65										108
						TTC Phe										156
GAG Glu	CTT Leu	CTT Leu	CAG Gln	CAA Gln -35	ATG Met	GAG Glu	AGA Arg	GGC Gly	CTG Leu -30	AAA Lys	TCA Ser	GCA Ala	GAC Asp	CBB Xaa -25	MSG Xaa	204
						GGC Gly										252
						GGG Gly										300
						CTG Leu										348
	CAA Gln															357

# (2) INFORMATION FOR SEQ ID NO: 252:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 414 base pairs

(B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR

# (ii) MOLECULĖ TYPE: CDNA

# (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

#### (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 11..238

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 1..228 id R26618

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 283..397

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 96..210 id HUM528H09B

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 202..282

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 16..96 id HUM528H09B

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 283..411

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 110..238 id C18739

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 202..282

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 30..110 id C18739

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 235..411

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97 region 1..177 id R17985 est

(ix	) FEATURE:
( <del>+</del> ~	L'ENIURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement (2..70)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 97 region 9..77 id R40947

est

#### (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 274..336
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 3.9

seq AWLAQGSSSAGWG/LE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 252:

ATCAATTIK TGAATAGTTT CCATTTCAAA TATCTTGTTC TACTTGGTTC ATAAAATAGT GGTTTTCAAA CTGTAGAGCT CTGGACTTCT CACTTCTAGG GCAGAGGGAG CCTGAACAAG TGAGGCTCTG GGTTCCCCAT TCCTAATTAA ACCAATGGAA AGAAGGGGTC TAATAACAAA 180 CTACAGCAAC ACATTTTCA TTTCAGCTTC ACTGCTGTAT CTCCCAGTCT AACCCTAGCA 240 TCCAGAAGTG GCACAAAACC CCTCTGCTGG CTC ATG TGT GCA ACT GAG ACT GTC 294 Met Cys Ala Thr Glu Thr Val AGA GCA TGG CTA GCT CAG GGG TCC AGC TCT GCA GGG TGG GGG CTA GAG 342 Arg Ala Trp Leu Ala Gln Gly Ser Ser Ser Ala Gly Trp Gly Leu Glu -10 -5 AGG AAG CAG GGA GTA TCT GCA CAC AGG ATG CCC GCG CTC AGG TGG TTG 390 Arg Lys Gln Gly Val Ser Ala His Arg Met Pro Ala Leu Arg Trp Leu 5 CAG AAG TCA GTG CCA GGA BCC ATG 414

# (2) INFORMATION FOR SEQ ID NO: 253:

Gln Lys Ser Val Pro Gly Xaa Met

20

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 189 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain

```
(ix) FEATURE:
```

- (A) NAME/KEY: other
- (B) LOCATION: 124..153
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 90

region 25..54

id N91869

est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 124..153
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 90

region 5..34 id H53427

est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 124..153
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 90

region 19..48 id H88369

est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 124..153
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 90

region 26..55

id T79771 est

# (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 124..153
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 90

region 29..58

id H41152

est

# (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 46..183
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 3.9

seq AAAFCLKXXGANT/HP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 253:

AGAATTTCTC CCACTCTTCG AGCCTACAGC AGACATGTTA GGAGA ATG CTG CTT 51

-45

105

GCA ACA CAC CCA GAG ACG GTG GGG CAG GTG ACA CTG CGT GTG TRC CCG

189

Ala Thr His Pro Glu Thr Val Gly Gln Val Thr Leu Arg Val Xaa Pro

-35

GTG TCT CTG GAA GTG TCT ATA CAG ATG TGT GCT GCT GCT GCT GCT Val Ser Leu Glu Val Ser Ile Gln Met Cys Ala Ala Ala Ala Ala Ala -15

TTC TGC CTT AAA ATK WCT GGA GCC AAC ACC CAC CCA

Phe Cys Leu Lys Xaa Xaa Gly Ala Asn Thr His Pro

-10 -5

(2) INFORMATION FOR SEQ ID NO: 254:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 300 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 149:.232

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 91..174 id AA081517

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 224..297

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 165..238

id AA081517

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 90..141

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 34..85

id AA081517

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 76..141

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 93 region 20..85

id N53273

									est							
	(:	ix)	(B) (C)	URE: NAMI LOCA I DEN	ATION NTIF:	N: 14	49 ION 1	<b>ЧЕТН</b>	ide:	ntity	y 97 91	135				
	(:	ix)	(B) (C)	URE: NAME LOCA IDEN OTHE	ATION NTIF:	N: co	omple ION N	AETH(	DD: h ider regi	olast ntity	n y 100 172					
			(B) (C) (D)	JRE: NAME LOCA I DEN OTHE	ATION NTIFI ER IN	I: 43 CATI	323 ION N MATIC	34 METHO DN:	DD: \ scoi seq	GLGC	9 SAQL(	QGGAX				
AGT	CTCT	GGG (	CGCGG	GCCA!	rg T	rgga	GGST	C CG	GCC	CGAG				GCG A		54
TCA Ser -60	GCA Ala	ACC Thr	CCA Pro	GCG Ala	CCS Pro -55	ASC Xaa	RGA Xaa	AGT Ser	CAG Gln	CGG Arg -50	TGC Cys	GGG Gly	GCA Ala	GAT Asp	GCT Ala -45	102
GGA Gly	AGT Ser	GCA Ala	GCC Ala	AGG Arg -40	ATT Ile	GTA Val	TTT Phe	CGG Arg	TGG Trp -35	GGC Gly	CGC Arg	GGC Gly	CGT Arg	CGC Arg -30	GGA Gly	150
GCC Ala	AGA Arg	TCA Ser	CCT Pro -25	GAG Glu	GGA Gly	AGC Ser	GGG Gly	CAT His -20	CAC His	GGC Gly	CGT Arg	GCT Ala	AAC Asn -15	AGT Ser	GGA Gly	198
CTC Leu	GGA Gly	GGA Gly	GCC Ala	CAG Gln	CTT Leu	CAA Gln	GGC	GGG Gly	GCC Ala	TRG Xaa	GGT Gly	CGA Arg	GGA Gly	TCT Ser	ATG Met	246

GCG CCT CTT CGT GCC AGC GCT GGA CAA ACC CGA GAC GGA CCT ACT CAG

Ala Pro Leu Arg Ala Ser Ala Gly Gln Thr Arg Asp Gly Pro Thr Gln
5 10 15 20

294

300

# (2) INFORMATION FOR SEQ ID NO: 255:

CCA GGG

Pro Gly

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 151 base pairs(B) TYPE: NUCLEIC ACID(C) STRANDEDNESS: DOUBLE(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

# (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

#### (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 13..150

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 1..138 id T36282 est

#### (ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 46..150

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 1..105 id T08090 est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 46..150

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 1..105 id T08091 est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 72..150

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 1..79 id H56620 est

#### (ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 80..150

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 1..71 id AA027983

est

#### (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 2..52

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 3.9

#### seq PLAGLAAAALGRA/PP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 255:

A ATG CTG CGG CGC CCG CTG GCC GGG CTG GCT GCG GCC CTG GGC CGG

Met Leu Arg Arg Pro Leu Ala Gly Leu Ala Ala Ala Leu Gly Arg

-15

GCC CCA CCG GAC GGC TTG CTC TGC TCT TTA CCT GGG GTT GCT GTC GAG

Ala Pro Pro Asp Gly Leu Leu Cys Ser Leu Pro Gly Val Ala Val Glu

1

5

GAC CCT GTG CAA GAC TCG GCC GGT TTT TCT TTC TCC CTG ATG GAC AGA

145

Asp Pro Val Gln Asp Ser Ala Gly Phe Ser Phe Ser Leu Met Asp Arg

20
25
30

CCC AAG
Pro Lys

# (2) INFORMATION FOR SEQ ID NO: 256:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 217 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 3..214
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 99 region 14..225 id H08058

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 2..91
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 100 region 10..99 id R11727

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 59..109
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.8 seq GFVAALVAGGVAG/VS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 256:

- (2) INFORMATION FOR SEQ ID NO: 257:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 158 base pairs
    - (B) TYPE: NUCLEIC ACID
    - (C) STRANDEDNESS: DOUBLE
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: CDNA
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: other
    - (B) LOCATION: 39..155
    - (C) IDENTIFICATION METHOD: blastn
    - (D) OTHER INFORMATION: identity 98 region 1..117

id C01598 est

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 9..71
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.8

seq SMDLLTLLFQRRS/HQ

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 257:

AATCAAGT ATG ATT GTG TTT GAG GGT ATT TCC ATG GAT CTC CTT ACA

Met Ile Val Trp Phe Glu Gly Ile Ser Met Asp Leu Leu Thr

-20

-15

-10

CTG CTA TTC CAG AGG AGA AGC CAC CAG GTC ACT CAA CTC TTA GTA TCA

Leu Leu Phe Gln Arg Arg Ser His Gln Val Thr Gln Leu Leu Val Ser

-5 1 5

TCT ACT GGA AAC TGG CTA AGA CAG TAT TTA TGT GCT TCT CTC ACA ATA

Ser Thr Gly Asn Trp Leu Arg Gln Tyr Leu Cys Ala Ser Leu Thr Ile

10 20 25

GCA GGA AGA AGG Ala Gly Arg Arg 158

#### (2) INFORMATION FOR SEQ ID NO: 258:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 292 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: complement(192..269)
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 92 region 354..431 id N70088

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (3) LOCATION: complement(222..262)
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 95 region 399..439 id H30254 est
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: 143..202
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.8

seq ALDALMFPARRRA/AV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 258:

AAGCGGCTGT CCTCCCTCGC TTTTGGAGCT CCGACCTCAG CTTCGCCTGC GAGCTGGGTT 60

GTGTAAAGGC TGGTCATTTT GGGGCGCTTA GGGGTGGGTG CCGGGGGGCG CGCTTTCCCT 120

CGTGAAGGTC GCTCCAGGAG TC ATG CGT ACA TTC GTT CAT TTT GCT CTG GAC 172

Met Arg Thr Phe Val His Phe Ala Leu Asp

-20 -15

GCA CTG ATG TTC CCG GCT CGC CGC CGT GCC GCA GTC ACG AGG CTC TCC
Ala Leu Met Phe Pro Ala Arg Arg Arg Ala Ala Val Thr Arg Leu Ser
-10 -5 1 5

GAA CGC CTT TCA CTG TGT TTC TGT TTA CAT TCG CGT CTG CAA GAC CCG
Glu Arg Leu Ser Leu Cys Phe Cys Leu His Ser Arg Leu Gln Asp Pro
10 15 20

GCG GCG CGA CCG AGG CCC TCT TGG
Ala Ala Arg Pro Arg Pro Ser Trp
25 30

#### (2) INFORMATION FOR SEQ ID NO: 259:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 338 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 131..273
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 92 region 120..262 id R10063

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 35..101
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 100 region 26..92

region 26... id R10063

est

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (3) LOCATION: 103..149
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 100

region 93..139

id R10063

- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 275..312
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 97

region 266..303 id R10063 est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 131..273

(C) IDENTIFICATION METHOD: blastn
(D) OTHER INFORMATION: identity 92

region 130..272

id R12045

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 35..100

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 36..101 id R12045

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 103..149

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 103..149

id R12045

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 3..35

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 5..37 id R12045

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 131..273

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 92

region 125..267

id R12117

est

(ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 5..100

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 1..96

id R12117

est

(ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 103..149

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 98..144 id R12117

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 131..273

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 92

region 102..244

id T79499

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 28..102

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 93

region 1..75 id T79499

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 104..149

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 76..121 id T79499

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 275..312

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 248..285

id T79499

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 109..178

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 109..178

id H17371

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 275..332

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 283..340

id H17371

est

#### (ix) FEATURE:

(A)	NAME/KEY: other
(3)	LOCATION: 44106
(C)	IDENTIFICATION METHOD: blastn
(G)	OTHER INFORMATION: identity 95
	region 42

region 42..104 id H17371 est

#### (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 42..224

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 3.8

seq LVMTFLFRNGSLQ/EK

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 259:

AGCTTACAGT TCCTAACCCC GACCCTGCGC GCASCCGCAC T ATG GCA GCC CCG CCG Met Ala Ala Pro Pro -60									
			CTG CTG CGC AAG CGC Leu Leu Arg Lys Arg -45	104					
			TTC CGG AAC GGG GCT Phe Arg Asn Gly Ala -25	152					
			CTG GTC ATG ACC TTT Leu Val Met Thr Phe -10	200					
			TGG GCC ATA CTG CAG Trp Ala Ile Leu Gln 5	248					
			TTT GTG TTC ACC TAC Phe Val Phe Thr Tyr 20	296					
	CGT GCC CTG CAG Arg Ala Leu Gln 30		Gly Pro Gly	338					

# (2) INFORMATION FOR SEQ ID NO: 260:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 364 base pairs
  - (3) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain

	i)	.x)	(B) (C)	JRE: NAME LOCA I-DEN OTHE	TION TIFI	1: 44 CATI	115 ON N	ETHO	ider regi	olast ntity on 2	, 99 208	322				
	<b>(</b> )	.x) l	(B) (C)	JRE: NAME LOCA IDEN OTHE	TION	: 28 CATI	373 ON M	ETHO	ider regi	olast ntity on 4	, 97 151	498				
	(i	.x) !	(B) (C)	JRE: NAME LOCA IDEN OTHE	TION TIFI	: 12 CATI	81 ON M	.81 1ETHC	D: V	on Hegen in the second in the	8					
	()	(i) {	SEQUE	ENCE	DESC	RIPT	: NOI	: SE(	) ID	NO:	260:	:				
ACCO	CAGCI	rcc (	CGGA	AGTGO	CG CC	CCGGI	AGCC	GCC	GCCG	CGGG	CCG	AGTG:	rcc 1	rggto	SAAGAC	60
CTAC	STTCI	TG (	CCGG	AGACA	AA TI	CCAC	CTGC	A GAA	AGCA	CTTT	ACT	(AAA)	AGG A	ACTTO	GCCAGG	120
CTGC	SACA			GTT Val												169
				GAG Glu 1												217
				TCC Ser												265
				GTG Val												313
				TTC Phe							Val					361
GGG Gly																364

# (2) INFORMATION FOR SEQ ID NO: 261:

(i) SEQUENCE CHARACTERISTICS:

276 . (A) LENGTH: 433 base pairs (B) TYPE: NUCLEIC ACID (C) STRANDEDNESS: DOUBLE (D) TOPOLOGY: LINEAR (ii) MOLECULE TYPE: CDNA (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo Sapiens (F) TISSUE TYPE: Brain (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: complement (324..433) (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 100 region 253..362 id H93008 (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: complement (200..267) (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 92 region 423..490 id H93008 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: complement(159..205) (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 93 region 484..530 id H93008 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: complement(116..162) (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 91 region 526..572 id H93008 est (ix) FEATURE: (A) NAME/KEY: other (B) LOCATION: complement(259..299) (C) IDENTIFICATION METHOD: blastn (D) OTHER INFORMATION: identity 97 region 390..430 id H93008 est (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (52..83)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 90

211

region 602..633 id H93008 est

#### (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 67..243

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 1..177 id AA146840 est

# (ix) FEATURE:

(A) NAME/KEY: other(B) LOCATION: 332..417

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 269..354 id AA146840 est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 242..299

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 177..234 id AA146840 est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 299..334

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 235..270 id AA146840

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 85..299

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 1..215 id AA036893

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 299..412

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 216..329

id AA036893

est

#### (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 98..243

(C) IDENTIFICATION METHOD: blastn
(D) OTHER INFORMATION: identity 98
region 1..146

id T49176 est

#### (ix) FEATURE:

- (A) NAME/KEY: other (B) LOCATION: 242..299
- (C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 146..203 id T49176 est

#### (ix) FEATURE:

- (A) NAME/KEY: other (B) LOCATION: 344..396
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 98

region 250..302 id T49176

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (3) LOCATION: 299..349
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 96

region 204..254 id T49176 est

### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 19..243
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 100

region 1..225 id H01262 est

#### (ix) FEATURE:

- (A) NAME/KEY: other
  (B) LOCATION: 242..296
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 100

region 225..279

id H01262

est

### (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 17..232
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 3.8

seq LMGLALAVYKCQS/MG

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 261:

ACTO	CAAAC	CAG A	ATTC(		eu Pi			et Al		AT AC	
	TCC Ser										100
	CAG Gln										148
	AGC Ser										196
	GGT Gly										244
	ACA Thr										292
	GAG Glu										340
	GGT Gly										388
	GCT Ala										433

# (2) INFORMATION FOR SEQ ID NO: 262:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 370 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 2..250
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 99 region 14..262

id N33874

(ix). FEATURE:

(A) NAME/KEY: other (B) LOCATION: 78..270

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 1..193 id H01141 est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 283..349

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 207..273 id H01141

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 284..366

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 92

region 320..402 id AA023741

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 74..270

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 2..198 id R27699

est

(ix) FEATURE:

(A) NAME/KEY: other (B) LOCATION: 320..349

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 93

region 253..282

id R27699

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement (320..366)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 282..328

id N33481

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement(283..322)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 327..366

id N33481

est

(ix)	FEATURE:
1	

- (A) NAME/KEY: other
- (B) LOCATION: complement (235..270)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 97

region 379..414 id N33481

est

#### (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 65..217
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 3.8

seq NVLFVAGLAFVIG/LE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 262:

ACGACTCAGC TTCCCACCCT	' GGGCTTTCCG	AGGTGCTTTC	GCCGCTGTCC	CCACCACTGC	60
-----------------------	--------------	------------	------------	------------	----

AGCC ATG ATC TCC TTA ACG GAC ACG CAG AAA ATT GGA ATG GGA TTA ACA

Met Ile Ser Leu Thr Asp Thr Gln Lys Ile Gly Met Gly Leu Thr

-50

-45

GGA TTT GGA GTG TTT TTC CTG TTC TTT GGA ATG ATT CTC TTT TTT GAC

Gly Phe Gly Val Phe Phe Leu Phe Phe Gly Met Ile Leu Phe Phe Asp

-35

-30

-25

AAA GCA CTA CTG GCT ATT GGA AAT GTT TTA TTT GTA GCC GGC TTG GCT
Lys Ala Leu Leu Ala Ile Gly Asn Val Leu Phe Val Ala Gly Leu Ala
-20 -15 -5

TTT GTA ATT GGT TTA GAA AGA ACA TTC AGA TTC TTC TTC CAA AAA CAT
Phe Val Ile Gly Leu Glu Arg Thr Phe Arg Phe Phe Gln Lys His

AAA ATG AAA GCT ACA GGT TTT TTT CTG GGT GTA TTT GTA GTC CTT
Lys Met Lys Ala Thr Gly Phe Phe Leu Gly Gly Val Phe Val Val Leu

15
20
25

ATT GGT TGG CCT TTG ATA GGC ATG ATC TTC GAA ATT TAT GGA TTT TTT

Ile Gly Trp Pro Leu Ile Gly Met Ile Phe Glu Ile Tyr Gly Phe Phe

30

40

CTC TTG TTC AGG GGC TTA GGG
Leu Leu Phe Arg Gly Leu Gly
'45

50

#### (2) INFORMATION FOR SEQ ID NO: 263:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 249 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: CDNA

#### (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

#### (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 112..249

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 153..290

id AA010288

est

#### (ix) FEATURE:

(A) NAME/KEY: other
(B) LOCATION: 112..218

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 96

region 101..207

id R26319

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 208..247

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 92

region 198..237

id R26319

est

### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 110..249

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 95

region 24..163

id W69087

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 112..247

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 103..238

id H01791

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: complement(112..217)

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 287..392

id AA146617

est

#### (ix) FEATURE:

		(B) (C)	NAME LOCA I DEN OTHE	TION TIFI	: 91 CATI	18 ON M	19 1etho	D: V	e 3.						
	(xi)	SEQU	ENCE	DESC	RIPT	CION:	SEC	Q ID	NO:	263:					
AAA	AAAGCGA	AGGC	CGGCC	G GG	CGGG	GAAG	G GGI	AAAT(	GCG	AGG	CAGG	AGT (	GCGG	GGAGG	60
GAG:	rggtcct ·	TAGC	TGAAT	rg co	CCT	GCGT					Gly			A AGG Arg	114
	CTG GT Leu Va														162
	ATG CT Met Le														210
	GAC CC Asp Pr														249
(2)	INFORM	SEQUEI (A) (B) (C)		HARA TH: NU	CTEF 324 CLEI	RISTI base CC AC	CS: pai								
	(ii)	MOLE	CULE	TYPE	: CI	ANG									
	(vi)		INAL ORGA TISS	NISM	l: Ho		•	ens							
	(ix)	(B) (C)	URE: NAME LOCA IDEN OTHE	TION TIFI	: 52 CAT	217 ION N	1ETHC	ider regi	ntity	7 98 58	L <b>94</b>				
	(ix)	(3) (C)	URE: NAME LOCA I DEN	ATION NTIFI	: 17 CAT	732 [ON 1	<b>ИЕТН</b> О	ide: reg.	ntit	y 98 190.	. 270				

	<b>(</b> i	ix). I	(B) (C)	NAME LOCA IDEN	E/KEY ATION NTIFI ER IN	1: 49 [CAT]	912 ION N	etho	D: / scoi	re 3.	6		atri: GQ/LF		
	()	<i) !<="" td=""><td>SEQUE</td><td>ENCE</td><td>DESC</td><td>CRIPT</td><td>rion:</td><td>: SE(</td><td>Q ID</td><td>NO:</td><td>264</td><td>:</td><td></td><td></td><td></td></i)>	SEQUE	ENCE	DESC	CRIPT	rion:	: SE(	Q ID	NO:	264	:			
AAG	GAGC'	TTC (	GCCG(	CGGC	CT GO	CTCC	SCCCi	A GC	CGGG	STCG	GTG	GCCG		TCG Ser	57
					TTC Phe										105
					GGG Gly										153
					CAG Gln 15										201
					GCT Ala										249
					CTA Leu										297
					CCA Pro										324
(2)		i) S1	EQUE	NCE (	SEQ CHARA	ACTE	RIST	cs:							
			(B) (C)	TYPE STRA	STH: E: NO ANDED OLOGY	ICLE I	C AC	CID DUBLE							
	( :	ii) 1	MOLE	CULE	TYP	E: CI	ANC								
	(1	vi) (	(A)	ORG	SOUI ANISM C SUE	1: Ho			ens						
	(:	ix)	(B) (C)	NAMI LOCA I DEI	E/KEY ATION NTIF: ER IN	N: CO	omple ION N	HTEN	DD: I ide: reg		in 796	226			

est

ŧ	(ix)	FEATURE	:
- 1		routone	٠

- (A) NAME/KEY: other
- (B) LOCATION: complement (43..156)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 98

region 73..186

id T23528

est

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: complement(6..156)
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 96

region 69..219

id R50519

est

#### (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 86..133
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 3.6

seq LAVLLVLFTLNIL/KS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 265:

# ACTGTATAAT RTGTGTATAT KAAAATGTAA TTGATTTCAG YYGAAAGTAT TTTAAAGCTG 60

ATAAATAGCA TTAGGGTTCT TTGCA ATG TGG TAT CTA GCT GTA TTA TTG GTT

Met Trp Tyr Leu Ala Val Leu Leu Val

-15
-10

TTA TTT ACT TTA AAC ATT TTG AAA AGC TTA TAC TGG CAG CCT GGG
Leu Phe Thr Leu Asn Ile Leu Lys Ser Leu Tyr Trp Gln Pro Gly
-5
1
5

# (2) INFORMATION FOR SEQ ID NO: 266:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 370 base pairs
  - (B) TYPE: NUCLEIC ACID
  - (C) STRANDEDNESS: DOUBLE
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain

#### (ix) FEATURE:

- (A) NAME/KEY: other
- (B) LOCATION: 41..79
- (C) IDENTIFICATION METHOD: blastn
- (D) OTHER INFORMATION: identity 92

286

region 80..118 id T06923

(ix) FEATU	RE:	٠
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- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: 197..322
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 3.5

seq INSLLEXSSLSRC/LE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 266:

ACATAAAGGA CASACGAGTC CTAATTGACA ACATCTAGTC TTTCTGGATG TTAAAGAGGT 60 TGCCAGTGTA TGACAAAAGT AGAGTTAGTA AACTAATATA TTTTGTACAT TTTGTTTTAC 120 AAGTCCTAGG AAAGATTGTC TTCTGAAAAT TTGATGTCTT CTGGGTTGAW GGAGATGGGA 180 AGGGTTCTAG GCCAGA ATG TTC ACA TTT GGA AGA CTC TTT CAA ATT ATA ACT 232 Met Phe Thr Phe Gly Arg Leu Phe Gln Ile Ile Thr GTT GTT ACA TGT TTG CAG TTT ATT CAA GAC TGC TGT ATA CAT AGT AGA 280 Val Val Thr Cys Leu Gln Phe Ile Gln Asp Cys Cys Ile His Ser Arg -25 -20 CAA ATT AAC TCC TTA CTT GAR RCA TCT AGT CTA TCT AGA TGT TTA GAA 328 Gln Ile Asn Ser Leu Leu Glu Xaa Ser Ser Leu Ser Arg Cys Leu Glu GTG CCG ATG TAT GTY AAA TGT ATA GGT AGT AAA ATA CCA CTT 370 Val Pro Met Tyr Val Lys Cys Ile Gly Ser Lys Ile Pro Leu

#### (2) INFORMATION FOR SEQ ID NO: 267:

5

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 301 base pairs

10

- (B) TYPE: NUCLEIC ACID
- (C) STRANDEDNESS: DOUBLE
- (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: CDNA
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: other
  - (B) LOCATION: 53..297
  - (C) IDENTIFICATION METHOD: blastn
  - (D) OTHER INFORMATION: identity 96 region 31..275 id HUM414A03B est

(ix)	(B) LOC (C) IDE	E/KEY: othe ATION: 22 NTIFICATION ER INFORMAT	51   METHOD:   ION: ide   rec	entity 10 gion 13 HUM414A0	0		
(ix)	(B) LOC	E/KEY: othe ATION: 106. NTIFICATION ER INFORMAT	.256 METHOD: MON: ide req	entity 99 gion 84 H83476			
(ix)	(B) LOC (C) IDE	E/KEY: othe ATION: 48 NTIFICATION ER INFORMAT	105   METHOD:   ION: ide   req	entity 96 gion 27 H83476			
(ix)	(B) LOC (C) IDE	E/KEY: sig_ ATION: 116. NTIFICATION ER INFORMAT	.268 METHOD: CION: sco	ore 3.5	ne matrix		
(xi)	SEQUENCE	DESCRIPTIO	ON: SEQ II	O NO: 267	':		
AAAATCAAGG	CAGGGGAT	GG AGGCAAG	rgg gggtc	GCGCC TGC	AGCGGAG (	CRTCGGCCTC	60
CGGAGCCGC	GCTGCAGC	CC TGTATTG	AGC TGAGA	IGGCT CGA	AGCCTAAC	ATTCC ATG Met	118
		AGA TGT GO Arg Cys A:					166
		A CGA GGC AG D Arg Gly T		y Pro Glu			214
		T GTG GGA A				Trp Ile	262
		C CCC CAC C S Pro His P			Trp		301

WO 99/06552 288

(i) SEQUENCE CHARACTERISTICS:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 404 base pairs(B) TYPE: NUCLEIC ACID(C) STRANDEDNESS: DOUBLE

(D) TO	POLOGY: LINEAR	
(ii) MOLECUL	E TYPE: CDNA	
	L SOURCE: GANISM: Homo Sapiens SSUE TYPE: Brain	
(B) LO (C) ID	ME/KEY: other CATION: 261404 ENTIFICATION METHOD: fasta HER INFORMATION: identity 100 region 1144 id HSU16126 vrt	
(B) LO (C) ID	E:  ME/KEY: sig_peptide  CATION: 261353  ENTIFICATION METHOD: Von Heijne matrix  HER INFORMATION: score 11.3  seq LLLCLLWIGYSQG/TT	
(xi) SEQUENC	E DESCRIPTION: SEQ ID NO: 268:	
AGGATTTCTC CCGGATG	CTC TCCGACTAAC ATGGATGTCC CACCATTCCT TGCAGTGGAA	60
GGTTGTTCCT TGGCGCA	GTG AGTGAAGAAC ATGCAGCGAT TGCTAATGGG TTTGGGAAGC	120
GGAGACTCCT TCCTCTC	TCT ATGACCATGC CGTGATCGTG TCTGCGGTCA CCACTCGACG	180
CATCCTCATT TCTACCC	GAA CCCAGGAGCC GAACGCTAGA TCGGGGAAGT GGGTGCCGTG	240
CGTGTGGGCA CAGAAAC	CACC ATG AAG ATT ATT TTC CCG ATT CTA AGT AAT CCA Met Lys Ile Ile Phe Pro Ile Leu Ser Asn Pro -30 -25	293
	CC GTT AAA CTC CTG CTC TGT TTA CTG TGG ATT GGA or Val Lys Leu Leu Cys Leu Leu Trp Ile Gly -15 -10 -5	341
	CC ACA CAT GTA TTA AGA TTT GGT GGT ATT TTT GAA or Thr His Val Leu Arg Phe Gly Gly Ile Phe Glu 1 5 10	389
TAT GTG GAA TCT GG Tyr Val Glu Ser Gl 15		404
(2) INFORMATION FO	DR SEQ ID NO: 269:	

(A) LENGTH: 249 base pairs

(B) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

## (ii) MOLECULE TYPE: CDNA

## (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 52..250

(C) IDENTIFICATION METHOD: fasta

(D) OTHER INFORMATION: identity 99

region 2..200 id HS7B2

vrt

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 14..250

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 24..260 id R14271

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 14..250

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

region 25..261

id R18347

est

#### (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 14..233

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 97

region 43..262

id H10233

est

# (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 14..240

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 99

region 44..270 id HSC0IE021

est

## (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 42..250

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 98

290

region 1..209 id HSCZSC021 est

1	ix	) FEATURE	:
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(A) NAME/KEY: sig\_peptide

(B) LOCATION: 79..156

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 6.6

seq LFWLASGWTPAFA/YS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 269:

AAGTTCGCCC GTNTCCTGGC CTGACCCCCA CCAAGGCCCA TACCGCAGTA GGCTCCTCGG								60					
GCT	SCCC	CTC (	GGTT	GACA		GTC Val -25		 	 		 		111
						GCA Ala				-	 		159
						CGG Arg			 		 		207
						CAA Gln		 	 				249

## (2) INFORMATION FOR SEQ ID NO: 270:

#### (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 316 base pairs

(3) TYPE: NUCLEIC ACID

(C) STRANDEDNESS: DOUBLE

(D) TOPOLOGY: LINEAR

## (ii) MOLECULE TYPE: CDNA

# (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

## (ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 212..311

(C) IDENTIFICATION METHOD: fasta

(D) OTHER INFORMATION: identity 93 region 1..101

id HSSCOASN

## (ix) FEATURE:

(A) NAME/KEY: other
(3) LOCATION: 243..311

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 94 region 60..128

id AA135265

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 187..245

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 91

region 5..63 id AA135265

est

(ix) FEATURE:

(A) NAME/KEY: other

(B) LOCATION: 269..311

(C) IDENTIFICATION METHOD: blastn

(D) OTHER INFORMATION: identity 100

region 49..91 id R58602 est

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: 179..250

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.8

seq ATMVSGSSGLAXA/RL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 270:

AGAGTTATTA TCTGCSTSTC CGATAGGATG CCTCTTTGTC TTCACCTGCC ATTCCCGCTG 60 TTTCGTGAAG AATCCTCTGT AAAGGGAAAT TTGTTCAGGC GACTGCTGTG GCCACCCTCT 120 GCCTCCTCCG GCCTCTGCCC CTGGGAGGTC CCCGGGGGCC TGGGAGTGTC ATTGGCGT ATG ACC GCA ACC CTT GCC GCT GCC GCT GAC ATC GCT ACC ATG GTC TCC 226 Met Thr Ala Thr Leu Ala Ala Ala Ala Asp Ile Ala Thr Met Val Ser GGC AGC AGC GGC CTC GCC GNC GCC CGT CTC CTG TCG CGC AST TCC TCC 274 Gly Ser Ser Gly Leu Ala Xaa Ala Arg Leu Leu Ser Arg Xaa Ser Ser TGC CGC AGA ATG GAA TTC GGC ATT GTT CCT ACA CAG CCA CGG 316 Cys Arg Arg Met Glu Phe Gly Ile Val Pro Thr Gln Pro Arg 10 15

# (2) INFORMATION FOR SEQ ID NO: 271:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 37 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -14..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 10.8

seq LLLLGLCLGLSLC/VG

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 271:

Met Leu Leu Gly Leu Cys Leu Gly Leu Ser Leu Cys Val Gly
-10 -5

Ser Gln Glu Glu Ala Gln Ser Trp Gly His Ser Ser Glu Gln Asp Gly
5 10 15

Leu Arg Val Pro Arg 20

- (2) INFORMATION FOR SEQ ID NO: 272:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 79 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -26..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 10.8

seq VLLFFVLLGMSQA/GS

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 272:
- Met Glu Asn Gly Gly Ala Gly Thr Leu Gln Ile Arg Gln Val Leu Leu -25 -15
- Phe Phe Val Leu Gly Met Ser Gln Ala Gly Ser Glu Thr Gly Asn -10 -5 1 5
- Phe Leu Val Met Glu Glu Leu Gln Ser Gly Ser Phe Val Gly Asn Leu 10 15 20
- Ala Lys Thr Leu Gly Leu Glu Val Ser Glu Leu Ser Ser Arg Gly Ala 35

Arg Val Val Ser Asn Asp Asn Lys Glu Cys Leu Gln Leu Asp Thr
40 45 50

(2) INFORMATION FOR SEQ ID NO: 273:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 129 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: -126..-1

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 10

seq LKLLLFLSTELQA/SQ

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 273:

Met Arg Gly Pro Glu Pro Gly Pro Gln Pro Thr Met Glu Gly Asp Val

Leu Asp Thr Leu Glu Ala Leu Gly Tyr Lys Gly Pro Leu Leu Glu Glu -110 -105 -100 -95

Gln Ala Leu Thr Lys Ala Ala Glu Gly Gly Leu Ser Ser Pro Glu Phe
-90 -85 -80

Ser Glu Leu Cys Ile Trp Leu Gly Ser Gln Ile Lys Ser Leu Cys Asr.
-75
-70
-65

Leu Glu Glu Ser Ile Thr Ser Ala Gly Arg Asp Asp Leu Glu Ser Phe
-60 -55 -50

Gln Leu Glu Ile Ser Gly Phe Leu Lys Glu Met Ala Cys Pro Tyr Ser -45 -40 -35

Val Leu Ile Ser Gly Asp Ile Lys Asp Arg Leu Lys Lys Lys Glu Asp
-30 -25 -20 -15

Cys Leu Lys Leu Leu Phe Leu Ser Thr Glu Leu Gln Ala Ser Gln
-10 -5

Ile

- (2) INFORMATION FOR SEQ ID NO: 274:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 44 amino acids
    - (B) TYPE: AMINO ACID

- (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -26..-1
  - · (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 9.6

seq WLIALASWSWALC/RI

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 274:

Met Glu Lys Ser Trp Met Leu Trp Asn Phe Val Glu Arg Trp Leu Ile
-25 -15

Ala Leu Ala Ser Trp Ser Trp Ala Leu Cys Arg Ile Ser Leu Leu Pro

Leu Ile Val Thr Phe His Leu Tyr Gly Gly Ser Gly
10 15

- (2) INFORMATION FOR SEQ ID NO: 275:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 89 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -36..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 9.5

seq LGLLLLARHWCIA/GV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 275:

Met Gln Gln Thr Arg Thr Glu Ala Val Ala Gly Ala Phe Ser His Cys
-35
-25

Leu Gly Phe Cys Gly Met Arg Leu Gly Leu Leu Leu Leu Ala Arg His -20 -15 -10 -5

Trp Cys Ile Ala Gly Val Phe Pro Gln Lys Phe Asp Gly Asp Ser Ala

Tyr Val Gly Met Ser Asp Gly Asn Pro Glu Leu Leu Ser Thr Ser Gln
15 20 25

Thr Tyr Asn Gly Gln Ser Glu Asn Asn Glu Asp Tyr Glu Ile Pro Pro 30 40

Ile Thr Pro Pro Asn Leu Pro Glu Ala 45 50

- (2) INFORMATION FOR SEQ ID NO: 276:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 53 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -24..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 9.1

seq LVVFLLLPLASGP/QV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 276:

Met Glu Lys Gly Asn Ala Phe Leu Lys Asn Arg Leu Val Val Phe Leu
-20 -15 -10

Leu Leu Pro Leu Ala Ser Gly Pro Gln Val Lys Arg Lys Ser Glu Ile -5 1 5

Thr Lys Leu Ile Lys Ala Thr Arg Ile Ile Cys Leu Phe Asn Lys Phe 10 20

Ser Arg Gly Asn Gly 25

- (2) INFORMATION FOR SEQ ID NO: 277:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 79 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -24..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 9

seq LLMLIVFHAASMA/LQ

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 277:

Met Phe Pro Phe Asn Gln Ala Gly Leu Pro Thr Leu Leu Met Leu Ile
-20 -15 -10

Val Phe His Ala Ala Ser Met Ala Leu Gln Arg Leu Phe Leu Phe Ala
-5 1 5

Leu Val Trp His Ser Lys Pro Ser Gly Leu Met Thr Gly Lys Leu Glu
10 20

Ser Gln Ile Pro His Glu Lys Leu Thr His Ile Ser Val Met His Gly 25 30 35 40

Pro Leu Ser Ser His His Ser Tyr Thr His Ile His Leu Phe Leu
45 50 55

- (2) INFORMATION FOR SEQ ID NO: 278:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 99 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -76..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 8.8

seq SLLLWMSSLPSLG/EK

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 278:

Met Thr Ser Arg Ser Leu Arg Arg Cys Ser Cys Leu Arg Val Thr His
-75 -70 -65

Asn Lys Glu Ile Leu Ala Ser Thr Val Ser Leu Gly Val Glu Giy Tyr
-60 -55 -50 -45

Met Leu Gly Gly Ser Arg Ile Asn Ser Ser Asn Leu Asn Asp Gly
-40 -35 -30

Glu Glu Glu Cys Ser Pro Asp Ser Leu Leu Val Trp Lys Lys Ser -25 -20 -15

Leu Leu Trp Met Ser Ser Leu Pro Ser Leu Gly Glu Lys Tyr Phe
-10 -5

Lys Arg Ile Leu Arg Trp Arg Glu His Trp Lys Ser Ser Gly Pro Ile 5 10 15 20

Pro Leu Trp

- (2) INFORMATION FOR SEQ ID NO: 279:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 68 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -53..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 8.8

seq ILLLLTVLPCIXM/GQ

PCT/IB98/01236

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 279:
- Met Trp Thr Ala Ser Ala Met Asp Phe Arg Thr Cys Ile Ala Ser Xaa -50 -45 -40
- Leu Pro Ala Leu Cys Tyr Val Gln Ala Cys Arg Ala Leu Met Ile Ala
  -35 -30 -25
- Ala Ser Val Leu Gly Leu Pro Ala Ile Leu Leu Leu Leu Thr Vai Leu
  -20 -15 -10
- Pro Cys Ile Xaa Met Gly Gln Glu Pro Gly Val Ala Lys Tyr Arg Xaa -5 10

Ala Gln Leu Ala

15

- (2) INFORMATION FOR SEQ ID NO: 280:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 86 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:

. (A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: -45..-1

- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 8.5

seq FALLSLSHPTCQA/GA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 280:

Met Gly Pro Pro Pro Thr His Ile Lys Tyr Leu His Leu Asn Ile Tyr
-45 -35 -30

Cys Asn Gly Lys Ser Thr Ala Pro Gly Ile Arg Ser His Ser Leu Gly
-25
-20
-15

Phe Ala Leu Leu Ser Leu Ser His Pro Thr Cys Gln Ala Gly Ala Pro
-10 -5 1

Val Arg Val Gly Arg Met Leu Ser His Leu Tyr Thr Cys Gly Trp Tyr 20 25 30 35

Asp His Asn Pro His Gly

- (2) INFORMATION FOR SEQ ID NO: 281:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 19 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -16..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 8.5

seg LLTFLAFTTLLFA/PP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 281:

Met Phe Cys Leu Leu Thr Phe Leu Ala Phe Thr Thr Leu Leu Phe Ala

Pro Pro Trp

(2) INFORMATION FOR SEQ ID NO: 282:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 80 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: -29..-1

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 8.4

seq LKCLLAVLSSLFA/AI

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 282:

Met His Cys Gly Ser Thr Pro Gly Leu Cys Pro Cys Trp Val Pro Phe
-25 -20 -15

Leu Lys Cys Leu Leu Ala Val Leu Ser Ser Leu Phe Ala Ala Ile Ser -10 -5 1

Val Asp Arg Leu Tyr Leu Ser Phe Cys Ser Asn Cys Ser Glu Ile Tyr
5 10 15

Leu Trp Pro Pro Ser Phe Pro Ala Pro Pro Ser Pro Val Val Leu Leu 20 25 30 35

Val Phe Leu Cys Pro His Gly Thr Ser Leu Ser Phe Leu Lys Leu Pro
40 45 50

- (2) INFORMATION FOR SEQ ID NO: 283:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 23 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -16..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 8.3

#### seq VCSALLLLGIVSS/KP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 283:

Met Asn Leu Val Cys Ser Ala Leu Leu Leu Gly Ile Val Ser Ser
-15 -10 -5

Lys Pro Tyr Met Arg Lys Arg

- (2) INFORMATION FOR SEQ ID NO: 284:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 47 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -35..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 8.3

seq AAMLIGLLAWLQT/VP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 284:

Met Ser Val Leu Asp Asp Arg Gln Arg Asp Ile Leu Val Val Gln Lys
-35
-20
-20

Arg His Ser Ser Leu Glu Ala Ala Met Leu Ile Gly Leu Leu Ala Trp
-15 -10 -5

Leu Gln Thr Val Pro Ala His Gly Cys Gln Phe Leu Pro Ile Arg 1 5 10

- (2) INFORMATION FOR SEQ ID NO: 285:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 38 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: -20..-1

- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 8.3 seq LLIICHYLPLSLC/IP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 285:

Met Gly Val Asn Gly Arg Arg Leu Leu Ile Ile Cys His Tyr Leu Pro -20 -15 -10 -5

Leu Ser Leu Cys Ile Pro Ile Pro Ser His Ile Asn Ser Leu Pro Arg  $1 \hspace{1.5cm} 5 \hspace{1.5cm} 10$ 

Asn Thr Pro Pro Val Arg 15

- (2) INFORMATION FOR SEQ ID NO: 286:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 43 amino acids
      - (B) TYPE: AMINO ACID
      - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -31..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 8.2 seq LECLLLYLAESSG/LR
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 286:

Met Lys Leu Arg Glu Cys Pro Ala Leu Arg Trp Ser Gln Leu Ser Gln -30 -25 -20

His Lys Leu Glu Cys Leu Leu Leu Tyr Leu Ala Glu Ser Ser Gly Leu
-15 -5 1

Arg Thr Gly Asn Val Gly Val Leu His Pro Arg
5 10

- (2) INFORMATION FOR SEQ ID NO: 287:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 136 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -109..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 8.2

seq LLRLPQLPPXCSA/GE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 287:

Met Asp Pro Arg Gly Ile Leu Lys Ala Phe Pro Lys Arg Gln Lys Ile
-105 -100 -95

His Ala Asp Ala Ser Ser Lys Val Leu Ala Lys Ile Pro Arg Arg Glu -90 -85 -80

Glu Gly Glu Glu Ala Glu Glu Trp Leu Ser Ser Leu Arg Ala His Val
-75 -70 -65

Val Arg Thr Gly Ile Gly Arg Ala Arg Ala Glu Leu Phe Glu Lys Gln
-60 -55 -50

Ile Val Gln His Gly Gly Gln Leu Cys Pro Ala Gln Gly Pro Gly Val -45 -40 -35 -30

Thr His Ile Val Val Asp Glu Gly Met Asp Tyr Glu Arg Ala Leu Arg
-25
-20
-15

Leu Leu Arg Leu Pro Gln Leu Pro Pro Xaa Cys Ser Ala Gly Glu Val

Ser Leu Ala Glu Leu Val Pro Ser Gly Glu Glu Ala Gly Gly Cys Ser 5 10 15

Trp Ile Gln His Leu His Pro Ser 20 25

- (2) INFORMATION FOR SEQ ID NO: 288:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 26 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -21..-1

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 8

seg LFLVAVLVKVAEA/RK

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 288:

Met Phe Trp Lys Leu Ser Leu Ser Leu Phe Leu Val Ala Val Leu Val -20 -15 -10

Lys Val Ala Glu Ala Arg Lys Asn Arg Ser
-5

- (2) INFORMATION FOR SEQ ID NO: 289:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 26 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -18..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7.9

seq LFSLLVLQSMATG/AT

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 289:

Thr Gly Ala Thr Phe Pro Glu Glu Ala Pro
1 5

- (2) INFORMATION FOR SEQ ID NO: 290:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 70 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide

- (B) LOCATION: -18..-1
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 7.9

seq LFSLLVLQSMATG/AT

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 290:

Met Ala Phe Leu Gly Leu Phe Ser Leu Leu Val Leu Gln Ser Met Ala
-15 -10 -5

Thr Gly Ala Thr Phe Pro Glu Glu Ala Ile Ala Asp Leu Ser Val Asn
1 5 10

Met Tyr Asn Arg Leu Arg Ala Val Gly Ser Trp Arg Arg Glu Gly Ala 15 20 25 30

Ser Arg Gln Ile Ala Ser Cys Leu Pro Ala Phe Leu Leu His Leu Pro 35 40 45

Leu Thr His Thr His Gly 50

- (2) INFORMATION FOR SEQ ID NO: 291:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 103 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -55..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7.8

seq ALLVALLFTLIHR/RR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 291:

Met Ser Phe Ser Leu Asn Phe Thr Leu Pro Ala Asn Thr Thr Ser Ser -55 -45 -45

Pro Val Thr Gly Gly Lys Glu Thr Asp Cys Gly Pro Ser Leu Gly Leu
-35
-30
-25

Ala Ala Gly Ile Pro Leu Leu Val Ala Thr Ala Leu Leu Val Ala Leu
-20
-15
-10

Leu Phe Thr Leu Ile His Arg Arg Arg Ser Ser Ile Glu Ala Met Glu
-5 5

Glu Ser Asp Arg Pro Cys Glu Ile Ser Glu Ile Asp Asp Asn Pro Lys

10 15 20 25

Ile Ser Glu Asn Pro Arg Arg Ser Pro Thr His Glu Lys Asn Thr Met 30 35 40

Gly Ala Gln Glu Ala Arg Trp
45

- (2) INFORMATION FOR SEQ ID NO: 292:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 82 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -80..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7.7

seq LVLFLSLALLVTP/TS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 292:

Met Ser Thr Trp Tyr Leu Ala Leu Asn Lys Ser Tyr Lys Asn Lys Asp
-80 -75 -70 -65

Ser Val Arg Ile Tyr Leu Ser Leu Cys Thr Val Ser Ile Lys Phe Thr
-60 -55 -50

Tyr Phe His Asp Ile Gln Thr Asn Cys Leu Thr Thr Trp Lys His Ser -45 -40 -35

Arg Cys Arg Phe Tyr Trp Ala Phe Gly Gly Ser Ile Leu Gln His Ser
-30 -25 -20

Val Asp Pro Leu Val Leu Phe Leu Ser Leu Ala Leu Leu Val Thr Pro
-15 -10 -5

Thr Ser

- (2) INFORMATION FOR SEQ ID NO: 293:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 43 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -19..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 7.7

seq LQLLCCIFTLVLQ/HY

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 293:

Met Ala Ile Gly Ile Ser Leu Gln Leu Leu Cys Cys Ile Phe Thr Leu
-15 -10 -5

Val Leu Gln His Tyr Leu Leu Gly Ser His Pro Tyr Ile Thr Cys Ile 1 5 10

His Ser Gln Leu Leu Leu Asp Ile Gln Gln 15 20

- (2) INFORMATION FOR SEQ ID NO: 294:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 74 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -19..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7.6

seq LLNLLLLSLFAGL/DP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 294:

Met Gln Ala Thr Ser Asn Leu Leu Asn Leu Leu Leu Leu Ser Leu Phe

Ala Gly Leu Asp Pro Ser Lys Asn Lys Lys Arg Gly Ser Ser Phe Ser  $1 \hspace{1cm} 5 \hspace{1cm} 10$ 

Phe Lys Phe Pro Leu Leu Asp Asp Thr Pro Phe Leu Xaa Ser Arg Ile 15 20 25

Glu Asn Ser Ala Thr His His Leu His Tyr Gly Leu Asn Met Ile Leu 30 40 45

Trp Val Asn Trp Lys Pro Lys Leu Thr Leu
50
55

- (2) INFORMATION FOR SEQ ID NO: 295:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 34 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -31..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7.6

seq VTLLCGWPGSHWC/AP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 295:

Met Met Lys Trp Lys Pro Glu Asp Leu Gly Ser Val Pro Cys Glu Ala
-30 -25 -20

Phe Ser Val Thr Leu Leu Cys Gly Trp Pro Gly Ser His Trp Cys Ala
-15 -5 1

Pro Pro

- (2) INFORMATION FOR SEQ ID NO: 296:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 78 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -19..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7.6

seq LLNLLLLSLFAGL/DP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 296:

Met Gln Ala Thr Ser Asn Leu Leu Asn Leu Leu Leu Leu Ser Leu Phe
-15 -10 -5

Ala Gly Leu Asp Pro Ser Lys Thr Gln Ile Ser Pro Lys Glu Gly Trp

1 5 10

Gln Val Tyr Ser Ser Ala Gln Asp Pro Asp Gly Arg Cys Ile Cys Thr 15 20 25

Val Val Ala Pro Glu Gln Asn Leu Cys Ser Arg Asp Ala Lys Ser Arg 30 40 45

Gln Leu Arg Gln Leu Leu Glu Lys Val Gln Asn Met Ser Arg 50 55

## (2) INFORMATION FOR SEQ ID NO: 297:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 80 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -48..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 7.5

seq FVILLLFIFTVVS/LV

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 297:
- Met Ala Ser Ser His Trp Asn Glu Thr Thr Thr Ser Val Tyr Gln Tyr
  -45 -40 -35
- Leu Gly Phe Gln Val Gln Lys Ile Tyr Pro Phe His Asp Asn Trp Asn -30 -25 -20
- Thr Ala Cys Phe Val Ile Leu Leu Phe Ile Phe Thr Val Val Ser
- Leu Val Val Leu Ala Phe Leu Tyr Glu Val Leu Asp Cys Cys Cys 1 5 10 15
- Val Lys Asn Lys Thr Val Lys Asp Leu Lys Ser Glu Pro Asn Pro Arg
  20 25 30
- (2) INFORMATION FOR SEQ ID NO: 298:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 35 amino acids

(B) TYPE: AMINO ACID
(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -33..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 7.4

seq ITCCVLLLLNCSG/VW

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 298:

Met Leu Trp Phe Ser Gly Val Gly Ala Leu Ala Glu Arg Tyr Cys Arg
-30 -25 -20

Arg Ser Pro Gly Ile Thr Cys Cys Val Leu Leu Leu Leu Asn Cys Ser
-15 -10 -5

Gly Val Trp 1

- (2) INFORMATION FOR SEQ ID NO: 299:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 31 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -25..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7.3

seq LIFFLNVTQLVRG/RG

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 299:

Met Leu Phe Leu Gln Met Gly Lys Gln Ser Trp Thr Leu Ile Phe Phe -25 -10 -15

Leu Asn Val Thr Gln Leu Val Arg Gly Arg Gly Pro Gly Gly Arg -5 1

(2) INFORMATION FOR SEQ ID NO: 300:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 36 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -27..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 7.2

seq LLLGLCSPPXXSL/AS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 300:

Met Glu Leu Arg Xaa Xaa Pro Pro Gly Gly Arg Glu Val Gln Leu Leu
-25 -20 -15

Leu Gly Leu Cys Ser Pro Pro Xaa Xaa Ser Leu Ala Ser Phe Pro Lys
-10 -5 1 5

Ala Ala Gln Met

- (2) INFORMATION FOR SEQ ID NO: 301:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 24 amino acids
    - (3) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -14..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7

seq LWSLLSSSGSHFG/IP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 301:

Met Leu Trp Ser Leu Leu Ser Ser Ser Gly Ser His Phe Gly Ile Pro

His His Thr Phe Pro Gln Glu Gly

(2) INFORMATION FOR SEQ ID NO: 302:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 112 amino acids
- (B) TYPE: AMINO ACID (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -52..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 7

seq SVWLCLLCYFAFP/FQ

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 302:

Met Asp Ile Ser Gly Leu Ile Pro Gly Leu Val Ser Thr Phe Ile Leu

Leu Ser Xaa Ser Asp His Tyr Gly Arg Lys Phe Pro Met Ile Leu Ser

Ser Val Gly Ala Leu Ala Thr Ser Val Trp Leu Cys Leu Leu Cys Tyr

Phe Ala Phe Pro Phe Gln Leu Leu Ile Ala Ser Thr Phe Ile Gly Ala 1

Phe Xaa Gly Asn Tyr Thr Thr Phe Trp Gly Ala Cys Phe Ala Tyr Ile

Val Asp Gln Cys Lys Glu Xaa Xaa Gln Lys Thr Ile Arg Ile Ala Ile

Ile Asp Phe Leu Leu Gly Leu Val Thr Gly Leu Thr Val Leu Ser Ser 50

- (2) INFORMATION FOR SEQ ID NO: 303:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 36 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

- (B) LOCATION: -30..-1
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 7

seq LFVILLITSLIFC/SL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 303:

Met Xaa Val Phe Phe Ser Lys Asn Arg Phe Glu Met Tyr Phe Ser Leu
-30 -25 -20 -15

Leu Leu Phe Val Ile Leu Leu Ile Thr Ser Leu Ile Phe Cys Ser Leu -10 -5 1

Tyr Val Ala Arg

- (2) INFORMATION FOR SEQ ID NO: 304:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 34 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -23..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 6.9

seq SLSLLASHHSVSC/SN

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 304:

Met Pro Val Pro Ala Cys Trp Ile Ser Ser Leu Ser Leu Leu Ala

Ser His His Ser Val Ser Cys Ser Asn Ile Phe Leu Asn Phe Asn Pro
-5 1 5

Asp Arg

- (2) INFORMATION FOR SEQ ID NO: 305:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 59 amino acids
    - (B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig peptide
  - (B) LOCATION: -21..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 6.9

seq LLACGSLLPGLWQ/HL

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 305:
- Met Cys Pro Val Phe Ser Lys Gln Leu Leu Ala Cys Gly Ser Leu Leu
  -20 -15 -10
- Pro Gly Leu Trp Gln His Leu Thr Ala Asn His Trp Pro Pro Phe Ser
  -5 1 5 10
- Xaa Phe Leu Cys Thr Val Cys Ser Gly Ser Ser Glu Gln Ile Ser Glu
  15 20 25
- Tyr Thr Ala Ser Ala Thr Pro Pro Leu Cys Leu 30 35
- (2) INFORMATION FOR SEQ ID NO: 306:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 128 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -76..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 6.9

seq LLPLSAWPPWAWH/HH

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 306:
- Met Ala Leu Thr Ile His Gly Glu Arg Met Arg Pro Asp Trp Glu Ser
  -75 -65
- Fro Trp Ile Thr Ser Ser Gln Ala Gln Ser Leu Ser Leu Gly Gly Ser
  -60 -55 -50 -45

Pro Ser Ser Arg Gly Pro Leu Val Pro Arg Gly Glu Tyr Leu Ala Ser

Cys Pro Glu Gly Val Arg Ser His Ser His Leu Leu Pro Arg Ser Leu
-25 -20 -15

Leu Pro Leu Ser Ala Trp Pro Pro Trp Ala Trp His His Gly Pro
-10 -5

Gly Thr Gln Ser Leu Val Gly Cys Leu Cys Ala Met Ser Pro Leu Leu 5 15 20

Pro Thr His Leu Ser Leu Pro Val Leu Glu Pro Ser Gly Thr Pro Ala 25 30 35

Leu Lys Asp Arg Arg Pro Cys Glu Val Gly Ile Pro Ile Pro Pro Arg
40 45 50

## (2) INFORMATION FOR SEQ ID NO: 307:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 95 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -92..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 6.8

seq ILIASSLPTLSHP/AP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 307:

Met Ala Ala Arg Phe Arg Cys Gly His Leu Cys Val Pro Glu Val Pro
-90 -85 -80

Arg Gly Pro Ala Ser His Ala Glu Gly Gly Gly Arg Leu Ser Arg

Lys Ala Ala His Gln Ala Gln Leu Cys Trp Arg Ala Gly Gly Asp Gly
-60 -55 -50 -50

Arg Gly Asn Phe Asn Pro Met Asn Phe Leu Val Ala Gly Thr Phe Ala
-40 -35 -30

Ser Ser Cys His Ser Pro Pro Leu Leu Trp Ser Leu Pro Pro Arg Ile
-25
-20
-15

Leu Ile Ala Ser Ser Leu Pro Thr Leu Ser His Pro Ala Pro Gly
-10 -5 1

(2) INFORMATION FOR SEQ ID NO: 308:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 87 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: -29..-1

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 6.8

seq VLSLICSCFYTQP/HP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 308:

Met Ala Ser Thr Ile Ser Ala Tyr Lys Glu Lys Met Lys Glu Leu Ser
-25 -20 -15

Val Leu Ser Leu Ile Cys Ser Cys Phe Tyr Thr Gln Pro His Pro Asn
-10 -5 1

Thr Val Tyr Gln Tyr Gly Asp Met Glu Val Lys Gln Leu Asp Lys Arg

Ala Ser Gly Gln Ser Phe Glu Val Ile Leu Lys Ser Pro Ser Asp Leu 20 25 30 35

Ser Pro Glu Ser Pro Met Leu Ser Ser Pro Pro Lys Lys Lys Asp Thr
40 45 50

Ser Leu Glu Glu Leu Gln Lys 55

(2) INFORMATION FOR SEQ ID NO: 309:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 120 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION:  $-11\overline{4}..-1$ 

WO 99/06552 PCT/IB98/01236

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 6.7 seq LIPMAILLGOTOS/NS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 309:

Met Leu Gln Val Tyr Gly Lys Pro Val Tyr Gln Gly His Arg Ser Thr -110 -105 -100

Leu Lys Lys Gly Pro Tyr Leu Arg Phe Asn Ser Pro Ser Pro Lys Ser
-95 -90 -85

Arg Pro Gln Arg Pro Lys Val Ile Glu Arg Val Lys Gly Thr Lys Val
-80 -75 -70

Lys Ser Ile Arg Thr Gln Thr Asp Phe Tyr Ala Thr Lys Pro Lys Lys
-65 -60 -55

Met Asp Ser Lys Met Lys His Ser Val Pro Val Leu Pro His Gly Asp
-50 -45 -40 -35

Gin Gln Tyr Leu Phe Ser Pro Ser Arg Glu Met Pro Thr Phe Ser Gly -30 -25 -20

Thr Leu Glu Gly His Leu Ile Pro Met Ala Ile Leu Leu Gly Gln Thr
-15 -10 -5

Gln Ser Asn Ser Asp Thr Met Pro

- (2) INFORMATION FOR SEQ ID NO: 310:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 127 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -118..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 6.7

seq LLFAKLFGHLTSA/RR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 310:

Met Ser Val Leu Glu Ile Ser Gly Met Ile Met Asn Arg Val Asn Ser -115 -105

His Ile Pro Gly Ile Gly Tyr Gln Ile Phe Gly Asn Ala Val Ser Leu
-100 -95 -90

Ile Leu Gly Leu Thr Pro Phe Val Phe Arg Leu Ser Gln Ala Thr Asp
-85 -80 -75

Leu Glu Gln Leu Thr Ala His Ser Ala Ser Glu Leu Tyr Val Ile Ala
-70 -65 -60 -55

Phe Gly Ser Asn Glu Asp Val Ile Val Leu Ser Met Val Ile Ile Ser -50 -45 -40

Phe Val Val Arg Val Ser Leu Val Trp Ile Phe Phe Leu Leu Cys
-35
-30
-25

Val Ala Glu Arg Thr Tyr Lys Gln Arg Leu Leu Phe Ala Lys Leu Phe
-20 -15 -10

Gly His Leu Thr Ser Ala Arg Arg Ala Arg Lys Ser Glu Val Pro
-5 5

## (2) INFORMATION FOR SEQ ID NO: 311:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 71 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -69..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 6.7 seq FFKLLLLGAMCSG/AR
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 311:

Met Cys Lys Gly Ile Lys Ala Gly Asp Thr Cys Glu Lys Leu Val Gly
-65 -60 -55

Tyr Ser Ala Val Tyr Arg Val Cys Phe Gly Met Ala Cys Phe Phe -50 -45 -40

Ile Phe Cys Leu Leu Thr Leu Lys Ile Asn Asn Ser Lys Ser Cys Arg
-35 -30 -25

Ala His Ile His Asn Gly Phe Trp Phe Phe Lys Leu Leu Leu Gly
-20 -15 -10

Ala Met Cys Ser Gly Ala Arg

(2) INFORMATION FOR SEQ ID NO: 312:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 114 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION:  $-10\overline{4}..-1$ 

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 6.6

seq HFSHVVWFHPTWA/QQ

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 312:

Met Ser Asp Ser Ala Gly Gly Arg Ala Gly Leu Arg Arg Tyr Pro Lys
-100 -95 -90

Leu Pro Val Trp Val Val Glu Asp His Gln Glu Val Leu Pro Phe Ile
-85 -80 -75

Tyr Arg Ala Ile Gly Ser Lys His Leu Pro Ala Ser Asn Val Ser Phe
-70 -65 -60

Leu His Phe Asp Ser His Pro Asp Leu Leu Ile Pro Val Asn Met Pro
-55 -50 -45

Ala Asp Thr Val Phe Asp Lys Glu Thr Leu Phe Gly Glu Leu Ser Ile
-40 -35 -30 -25

Glu Asn Trp Ile Met Pro Ala Val Tyr Ala Gly His Phe Ser His Val

Val Trp Phe His Pro Thr Trp Ala Gln Gln Ile Arg Glu Gly Arg His

His Phe

10

(2) INFORMATION FOR SEQ ID NO: 313:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 109 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

PCT/IB98/01236 WO 99/06552

- (ix) FEATURE:
  - (A) NAME/KEY: sig peptide
  - (B) LOCATION: -47..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 6.3

seq SSCVLLTALVALA/AY

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 313:

Met Ser Ser Cys Arg Gly Gln Lys Val Ala Gly Gly Leu Arg Val Val

Ser Pro Phe Pro Leu Cys Gln Pro Ala Gly Glu Pro Ser Arg Gly Lys -25

Met Arg Ser Ser Cys Val Leu Leu Thr Ala Leu Val Ala Leu Ala Ala

Tyr Tyr Val Tyr Ile Pro Leu Pro Gly Ser Val Ser Asp Pro Trp Lys

Leu Met Leu Leu Asp Ala Thr Phe Arg Gly Ala Xaa Xaa Xaa Ser Xaa

Leu Val Xaa Tyr Leu Gly Leu Ser Xaa His Leu Leu Ala Leu Xaa Xaa

Xaa Leu Phe Leu Leu Ala Lys Lys Ala Arg Gly Leu Leu

- (2) INFORMATION FOR SEQ ID NO: 314:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 72 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -42..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 6.1

seq DLAVALSLLPAWT/ES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 314:

Met Ile Ile Pro Phe Lys Ile Lys Asn Leu Gly Gly Arg Val Leu Leu -30 -40

Ser Gly Arg Glu Met Phe Pro Ala Ser Val Arg Ala Pro Asp Leu Ala

-25 -20 **-**15

Val Ala Leu Ser Leu Leu Pro Ala Trp Thr Glu Ser Pro Thr Arg Gly
-10 -5 1 5

Ser His Gln Ser Gln Ala Arg Ala His Ser Arg Ala Leu Arg Lys Gln 10 15 20

Ser Arg Asn Thr Arg Ser Pro Arg 25 30

- (2) INFORMATION FOR SEO ID NO: 315:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 60 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -53..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 6

seq ALILLLLAQKGPS/XF

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 315:
- Met Val Cys Ser Ala Pro Arg Lys Ile Val Val Arg Ala Phe Ile Thr
  -50 -45 -40
- Ile Ile Phe Ile Tyr Tyr Ala Ile Lys Lys Arg Ala Asn Glu Pro Ala
  -35
  -30
  -25
- Ala Tyr Leu Met Leu Lys Pro Glu Ala Leu Ile Leu Leu Leu Ala
  -20 -15 -10
- Gln Lys Gly Pro Ser Xaa Phe Leu Leu Val Trp Arg
  -5 5
- (2) INFORMATION FOR SEQ ID NO: 316:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 60 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -40..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5.9

seq VCSALCSLGEVRP/XE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 316:

Met Thr Glu Ser Ser Met Lys Lys Leu Ala Ser Thr Leu Leu Asp Ala
-40 -35 -30 -25

Ile Thr Asp Lys Asp Pro Leu Val Gln Glu Gln Val Cys Ser Ala Leu
-20 -15 -10

Cys Ser Leu Gly Glu Val Arg Pro Xaa Glu Thr Leu Arg Ala Cys Glu
-5 1 5

Glu Tyr Leu Arg Xaa Met Thr Ser Trp His Thr Arg
10 15 20

- (2) INFORMATION FOR SEQ ID NO: 317:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 59 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -43..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.9 seq VFLFHCTSGLSSC/KC
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 317:

Met Gln Glu Thr Asp Cys Asn Lys Arg Trp Gly Arg Gly Leu Gly Gly
-40 -35 -30

Leu Trp Ser Glu Thr Gly Arg Arg Phe His Cys Lys Ser Phe Val Phe
-25
-20
-15

Leu Phe His Cys Thr Ser Gly Leu Ser Ser Cys Lys Cys Ser Lys Lys
-10
-5
1
5

His Xaa Lys Tyr Cys Phe Cys Phe Val Ala Ser

Ser Pro Ala Phe Leu Ala Val Ala Gly Pro Gly Trp Ala Arg Pro Gly
-10 -5 1

Cys Xaa Leu Arg Thr Lys Tyr Asp Ser Gln Leu Ala Arg His Leu Leu 5 10

Gln Pro Gln Phe Pro Gly Leu Thr Leu Gly Thr Leu Val Gln Pro Ala 20 25 30 35

His Trp Gly Met Gly Gly Gly Thr Gly Gly Val Leu Gly Glu Gly Gly 40 45 50

Gly His Ser Tyr Ala Glu His Gly Thr Cys Leu Gln Ser Cys Ser Thr 55 60 65

Asp Val Leu Xaa His Val Leu Leu Ala

- (2) INFORMATION FOR SEQ ID NO: 320:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 125 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -18..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.6

seq WHFLASFFPRAGC/HG

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 320:

Met Leu Gln Met Leu Trp His Phe Leu Ala Ser Phe Phe Pro Arg Ala

Gly Cys His Gly Ser Arg Glu Gly Asp Asp Arg Glu Val Arg Gly Thr 1 5 10

Pro Ala Pro Ala Trp Arg Asp Gln Met Ala Ser Phe Leu Gly Lys Gln 15 20 25 30

Asp Gly Arg Ala Glu Ala Thr Glu Lys Arg Pro Thr Ile Leu Leu Val
35 40 45

Val Gly Pro Ala Glu Gln Phe Pro Lys Lys Ile Val Gln Ala Gly Asp 50 55 60

Lys Asp Leu Asp Gly Gln Leu Asp Phe Glu Glu Phe Val His Tyr Leu 65 70 75

(2) INFORMATION FOR SEQ ID NO: 318:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 45 amino acids

- (B) TYPE: AMINO ACID
- (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -17..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5.8

seq VPWLSSTVSCAQG/LR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 318:

Met Leu Leu Glu Val Pro Trp Leu Ser Ser Thr Val Ser Cys Ala Gln -15

Gly Leu Arg Leu Ala Gln His Arg Val Pro Phe Phe Tyr Ser Asn Val

Ser Leu Cys Lys Leu Leu Pro Ala Xaa Leu His Gly

- (2) INFORMATION FOR SEQ ID NO: 319:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 105 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -29..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.7

seq SPAFLAVAGPGWA/RP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 319:

Met Ser Gly Gly Arg Met Gln Ala Arg Cys Ser Gln Gln Ser Thr Trp -20 -25

Gln Asp His Glu Lys Lys Leu Arg Leu Val Phe Lys Ser Leu Asp Lys 80 85 90

Lys Asn Asp Gly Arg Ile Asp Ala Gln Glu Ile Met Gln 95 100 105

- (2) INFORMATION FOR SEQ ID NO: 321:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 37 amino acids
    - (B) TYPE: AMINO ACTD
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -20..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.6

seq SLVCLLAMGKGLG/SS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 321:

Met Tyr Ser His Pro Val Ser Ser Leu Val Cys Leu Leu Ala Met Gly -20 -15 -10 -5

Lys Gly Leu Gly Ser Ser Gln Ala Leu Val Gln Pro Asp Thr Trp Pro  $1 \hspace{1cm} 5 \hspace{1cm} 10$ 

His Thr Ser Pro Arg
15

- (2) INFORMATION FOR SEQ ID NO: 322:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 92 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -35..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.6

## seq FIFMEVLGSGAFS/EV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 322:

Met Gly Arg Lys Glu Glu Asp Asp Cys Ser Xaa Trp Lys Lys Gln Thr
-35 -25 -20

Thr Asn Ile Arg Lys Thr Phe Ile Phe Met Glu Val Leu Gly Ser Gly
-15 -10 -5

Ala Phe Ser Glu Val Phe Leu Val Lys Gln Arg Leu Thr Gly Lys Leu  $1 \hspace{1.5cm} 5 \hspace{1.5cm} 10$ 

Phe Ala Leu Lys Cys Ile Lys Lys Ser Pro Ala Phe Arg Asp Ser Ser 15 20 25

Leu Glu Asn Glu Ile Ala Val Leu Lys Lys Ile Lys His Glu Asn Ile 30 40 45

Val Thr Leu Glu Asp Ile Tyr Glu Ser Thr Gln Gly

- (2) INFORMATION FOR SEQ ID NO: 323:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 48 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -29..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.6

seq LLPNQSLFSLARA/VR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 323:

Met Met Ile Ala Val Phe Gly Asn Ala Asn Asp Arg Asn Val Leu Thr

Leu Leu Pro Asn Gln Ser Leu Phe Ser Leu Ala Arg Ala Val Arg Asn -10 -5 1

His Leu Leu Glu Glu Arg Arg Leu Thr Thr Tyr Gly Val Leu Cys
5 10 15

(2) INFORMATION FOR SEQ ID NO: 324:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 58 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -19..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5.6

seq LVVTAWFFGMCRS/KA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 324:

Met Phe Phe Glu Leu Pro Leu Val Val Thr Ala Trp Phe Phe Gly Met
-15 -10 -5

Cys Arg Ser Lys Ala Leu Leu Gly Asn Ala Arg Ser Ala Leu Cys Leu 1 5 10

Gln Thr Lys Ala Cys Ala Ser Ser Thr Gln Pro Asp Thr His Asn Glu 15 20 25

His His Pro Arg Asn Pro Cys Pro Tyr Leu 30 35

- (2) INFORMATION FOR SEQ ID NO: 325:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 54 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -44..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.5

seq FLLIVANVHFSQT/WV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 325:

Met Asn His Asn Ile Ile Ile Cys Val Met Tyr Ile Val Pro Phe Leu
-40 -35 -30

Met Thr Lys Cys Leu Tyr Phe Cys His Ser Cys Lys Arg Gly Ser Phe

Leu Leu Ile Val Ala Asn Val His Phe Ser Gln Thr Trp Val Phe Ser -10 -5 1

Gly Lys Pro Tyr Lys Gly
5

- (2) INFORMATION FOR SEQ ID NO: 326:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 53 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -21..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.5 seq LTGLCXCCLQALG/LA
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 326:

Met Ser Cys Gly Ser Ala Ala Ser Leu Thr Gly Leu Cys Xaa Cys Cys -20 -15 -10

Leu Gln Ala Leu Gly Leu Ala Trp Arg Arg Gly Leu Thr Gly Pro
-5 1 5 10

Gly Leu Pro Pro Val Leu Gln Ile Cys Cys Pro Arg Ser Leu Arg Gly
15 20 25

Val Thr Ala Pro Thr 30

- (2) INFORMATION FOR SEQ ID NO: 327:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 68 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: -46..-1

- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 5.4

seq VLFFVGLITNGLA/MR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 327:

Met Gln Ala Val Asp Asn Leu Thr Ser Ala Pro Gly Asn Thr Ser Leu
-45 -35

Cys Thr Arg Asp Tyr Lys Ile Thr Gln Val Leu Phe Pro Leu Leu Tyr -30 -25 -20 -15

Thr Val Leu Phe Phe Val Gly Leu Ile Thr Asn Gly Leu Ala Met Arg

Ile Phe Phe Gln Ile Arg Ser Lys Ser Asn Phe Ile Ile Phe Leu Lys
5 10 15

Asn Thr Val Lys

- (2) INFORMATION FOR SEO ID NO: 328:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 28 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -20..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.3

seq LCSSCCSWGPAAG/AL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 328:

Met Ala Ala Met Xaa Leu Leu Cys Ser Ser Cys Cys Ser Trp Gly -20 -15 -10 -5

Pro Ala Ala Gly Ala Leu Gln Asn Pro Gln Arg Gly
1 5

- (2) INFORMATION FOR SEQ ID NO: 329:
  - (i) SEQUENCE CHARACTERISTICS:

WO 99/06552 PCT/IB98/01236

(A) LENGTH: 30 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -25..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5.3

seq SVVKVLSLRKAQA/QS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 329:

Met Asp Phe Ile Lys Asp Gln Ser Leu Ser His Arg Ser Val Val Lys
-25
-15
-10

Val Leu Ser Leu Arg Lys Ala Gln Ala Gln Ser Ile Leu Glu
-5 1 5

- (2) INFORMATION FOR SEQ ID NO: 330:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 75 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -28..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.3

seq RISCAFSLASSTA/RQ

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 330:

Met Thr Arg Pro Phe Trp Ala Ser Cys Ser Thr Trp Ala Thr Ser Arg
-25 -20 -15

Ile Ala Cys Cys Ala Thr His Arg Thr Ala Trp Ala Ser Arg Pro Gly
5 10 15 20

Pro Arg Arg Pro Trp Cys Cys Arg Tyr Ser Lys Pro Leu Thr Trp

25 30 35

Pro Val Arg Met Met Arg Arg Glu Gly Ser Xaa 40 45

- (2) INFORMATION FOR SEQ ID NO: 331:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 35 amino acids
    - (B) TYPE: AMINO ACID
      (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -16..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.3

seq CAVSLTTAAVAFG/DE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 331:

Met Lys Ser Cys Ala Val Ser Leu Thr Thr Ala Ala Val Ala Phe Gly

Asp Glu Ala Lys Lys Met Ala Glu Gly Lys Ala Ser Arg Glu Ser Glu 1 5 10 15

Glu Glu Thr

- (2) INFORMATION FOR SEQ ID NO: 332:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 39 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -21..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.2

seq LSLSLICLRMSLS/LY

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 332:

Met Ser Ile His Glu Cys Ala Cys Leu Ser Leu Ser Leu Ile Cys Leu
-20 -15 -10

Arg Met Ser Leu Ser Leu Tyr Pro Pro Pro Ala Ser Met Ile Leu Leu -5 5 10

Pro Gln Thr Trp Lys Pro Arg

- (2) INFORMATION FOR SEQ ID NO: 333:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 42 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -15..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.2

seq SGLSFLSVFSLWC/EP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 333:

Met Leu Ser Gly Leu Ser Phe Leu Ser Val Phe Ser Leu Trp Cys Glu -15 -5 1

Pro Thr Leu Pro Ala Leu Gly Asn Gly Ser Val Leu Gly Val Arg Xaa 5 10 15

Ser Ser Ser Ser Ala Gln Cys Ser Leu 20 25

- (2) INFORMATION FOR SEQ ID NO: 334:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 87 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:

PCT/IB98/01236 WO 99/06552

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: -85..-1
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 5.2

seq LYSILHFPFWVHG/RX

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 334:

Met Gly Leu Lys Asp Lys Ser Gln Ala Pro Ala Ser Gly Leu Gly Val

Leu Arg Gly Gln Arg Ser Gly Ser Phe Ile Ser Met Pro Ala Pro Ala

Ser Gly Gln Xaa Pro Glu Glu Ser Arg Ser Pro Ala Pro Pro Val Ala -45

Ser Arg Ser Gln Asn Arg Gly Tyr Arg Pro Trp His Gly Pro Leu Trp

Val His Gln Ser Val Arg Phe Gly Leu Tyr Ser Ile Leu His Phe Pro

Phe Trp Val His Gly Arg Xaa

## (2) INFORMATION FOR SEQ ID NO: 335:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 67 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -43..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5.2

seq PMQLLQVLSDVLA/EI

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 335:

Met Ser Asp Gln Ile Lys Phe Ile Met Asp Ser Leu Asn Lys Glu Pro -35

Phe Arg Lys Asn Tyr Asn Leu Ile Thr Phe Asp Ser Leu Glu Pro Met -20

Sin Leu Leu Gin Val Leu Ser Asp Val Leu Ala Glu Ile Asp Pro Lys

Val Arg Val Phe Ser Phe Phe Leu Met Gly Ser Arg Lys Pro Ile Ser 10 15 20

Pro Ser Trp

- (2) INFORMATION FOR SEQ ID NO: 336:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 110 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig peptide
    - (B) LOCATION: -104..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.1

seq SSVASLTATPSLA/SP

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 336:
- Met Ser Pro Ser Cys Leu His Pro Asp Leu Trp Ser Met Cys Leu Glu
  -100 -95 -90
- Val Pro Ser Phe Thr Ala Thr Asp Ser Val Asn Cys Gly Cys Leu
  -85
- Glu Leu Ala Thr Glu Pro Ala Arg Asn Ile Arg Ser Thr Thr Arg Ala
  - Ser Leu Leu Arg Cys Ser Ser Phe Thr Ser Thr Arg Asn Ser Thr Gly
    -55 -50 -45
  - Ile Ser Ala Leu Pro Pro Ala Ala Pro Met Ala Trp Pro Phe Ser Ala
    -40 -35 -30 -25
  - Ser Leu Ser Thr Leu Pro Val Pro Leu Thr His Ser Ser Val Ala Ser
  - Leu Thr Ala Thr Pro Ser Leu Ala Ser Pro Thr Arg Met Met -5 1 5
  - (2) INFORMATION FOR SEQ ID NO: 337:
    - (i) SEQUENCE CHARACTERISTICS:
      - (A) LENGTH: 24 amino acids
      - (B) TYPE: AMINO ACJD
      - (D) TOPOLOGY: LINEAR
    - (ii) MOLECULE TYPE: PROTEIN

- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -16..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5.1

seq SFHLLLDPSSTQS/SI

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 337:

Met Asp Leu Ser Phe His Leu Leu Asp Pro Ser Ser Thr Gln Ser
-15 -5

Ser Ile Leu Lys His Leu Pro Cys

- (2) INFORMATION FOR SEQ ID NO: 338:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 53 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -26..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5

seq VISVLILVGFGAC/IY

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 338:

Met Pro His Phe Leu Asp Trp Phe Val Xaa Val Tyr Leu Val Ile Ser

Val Leu Ile Leu Val Gly Phe Gly Ala Cys Ile Tyr Tyr Phe Glu Pro -10 -5

Gly Leu Gln Glu Ala His Lys Trp Arg Met Xaa Arg Pro Trp Thr

Ala Thr Ser Thr Gly 25

(2) INFORMATION FOR SEQ ID NO: 339:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 39 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -30..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5

seq IVGLLAQLEKINA/EP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 339:

Met Ser Lys Leu Lys Val Ile Pro Glu Lys Ser Leu Thr Asn Asn Ser
-30 -25 -20 -15

Arg Ile Val Gly Leu Leu Ala Gln Leu Glu Lys Ile Asn Ala Glu Pro
-10 -5 1

Ser Glu Ser Asp Thr Ser Arg
5

- (2) INFORMATION FOR SEQ ID NO: 340:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 43 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -23..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5

seq LIPAMAFLSCVRP/ES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 340:

Met Met Ser Ala Ser Arg Leu Ala Gly Thr Leu Ile Pro Ala Met Ala -20 -15 -10

Phe Leu Ser Cys Val Arg Pro Glu Ser Xaa Glu Pro Cys Val Glu Val -5 5

Val Pro Asn Ile Thr Tyr Gln Cys Met Glu Leu 10 15 20

- (2) INFORMATION FOR SEQ ID NO: 341:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 65 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -49..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.9

seq VTVCCXLVAFLFC/IL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 341:

Met Val Asp Gly Thr Gln Leu Arg Gly Leu Thr Arg Met Tyr Gln Val
-45 -40 -35

Pro Leu Xaa Leu Asp Arg Asp Glu Thr Leu Val Arg Leu Arg Phe Thr
-30 -25 -20

Met Val Ala Leu Val Thr Val Cys Cys Xaa Leu Val Ala Phe Leu Phe
-15
-10
-5

Cys Ile Leu Trp Ser Leu Leu Phe His Phe Lys Glu Thr Thr Ala Thr
1 5 10 15

Gly

- (2) INFORMATION FOR SEQ ID NO: 342:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 34 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (3) LOCATION: -26..-1

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.9 seq LISMLQMLAVIIT/NT

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 342:

Met Lys Gln Asn Phe Leu Val Leu Asn Ser Val Trp Tyr Leu Ile Ser
-25 -20 -15

Met Leu Gln Met Leu Ala Val Ile Ile Thr Asn Thr Thr Ile Thr Thr -10 -5 1 5

Ile Gly

- (2) INFORMATION FOR SEQ ID NO: 343:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 124 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -59..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.9

seq LVEMCLEVLGSSA/GD

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 343:

Met Glu Cys Gln Asn Ser Ser Leu Lys Lys Cys Leu Leu Val Glu Lys
-55
-50
-45

Ser Leu Val Lys Ala Ser Tyr Leu Ile Ala Phe Gln Thr Ala Ala Ser
-40 -35 -30

Lys Lys Pro Phe Ser Ile Ala Glu Glu Leu Ile Lys Pro Tyr Leu Val

Glu Met Cys Leu Glu Val Leu Gly Ser Ser Ala Gly Asp Lys Met Lys
-10 -5 1 5

Thr Ile Pro Leu Ser Asn Val Thr Ile Gln His Arg Ile Asp Glu Leu 10 15 20

Ser Ala Asp Ile Glu Asp Gln Leu Ile Gln Lys Val Arg Glu Ser Lys 25 30 35

Trp Phe Ala Leu Gln Ile Asp Glu Ser Ser Glu Ile Ser Asn Ile Thr 40 45 50

PCT/IB98/01236 WO 99/06552 338

Leu Leu Cys Tyr Ile Arg Phe Ile Asp Tyr Asp 60

- (2) INFORMATION FOR SEQ ID NO: 344:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 27 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -23..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.8

seq VMWLVALLEMCVC/KK

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 344:

Met His Ser Ser Ile Lys Thr Lys Gly Ser Val Met Trp Leu Val Ala

Leu Leu Glu Met Cys Val Cys Lys Ser Arg

- (2) INFORMATION FOR SEQ ID NO: 345:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 44 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -18..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.8

seq LEAISSLSSFVLG/RM

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 345:

Met Thr Val Leu Pro Leu Glu Ala Ile Ser Ser Leu Ser Ser Phe Val -15 -10 -5

Leu Gly Arg Met Asn Ser Arg Gly Ala Gly Lys Thr Gln Asn Leu Asp

Ala Ser Ser Leu Leu Leu Leu Cys Cys Leu Ile Leu 15 20 25

- (2) INFORMATION FOR SEQ ID NO: 346:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 125 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -30..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.8

seq ILFCVGAVGACTL/SV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 346:

Met Gly Thr Ala Ser Arg Ser Asn Ile Ala Arg His Leu Gln Thr Asn
-30
-25
-20
-15

Leu Ile Leu Phe Cys Val Gly Ala Val Gly Ala Cys Thr Leu Ser Val -10 -5 1

Thr Gln Pro Trp Tyr Leu Glu Val Asp Tyr Thr His Glu Ala Val Thr 5 10 15

Ile Lys Cys Thr Phe Ser Ala Thr Gly Cys Pro Ser Glu Gln Pro Thr 20 25 30

Cys Leu Trp Phe Arg Tyr Gly Ala His Gln Pro Glu Asn Leu Cys Leu 35 40 45 50

Asp Gly Cys Lys Ser Glu Ala Xaa Lys Phe Thr Val Arg Glu Ala Leu
55 60 65

Lys Glu Asn Gln Val Ser Leu Thr Val Asn Arg Val Thr Ser Asn Asp 70 75 80

Ser Ala Ile Tyr Ile Cys Gly Ile Ala Phe Pro Ser Val 85 90 95

- (2) INFORMATION FOR SEQ ID NO: 347:
  - (i) SEQUENCE CHARACTERISTICS:

WO 99/06552 PCT/IB98/01236

(A) LENGTH: 46 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

- (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig peptide
  - (B) LOCATION: -25..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.8

seq ALFYSVVVSTVSG/NE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 347:

Met Asn Ser Ser Lys Glu Glu Met Arg Glu Leu Ala Ala Leu Phe Tyr
-25
-10
-10

Ser Val Val Ser Thr Val Ser Gly Asn Glu Leu Lys Ser Met Ile
-5 1 5

Glu Gln Leu Ile Lys Thr Thr Lys Asp Asn His Ser Leu Arg
10 15 20

- (2) INFORMATION FOR SEQ ID NO: 348:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 81 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig peptide
    - (B) LOCATION: -52..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.7

seq LLAKALHLLKSSC/AP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 348:

Met Ser Gln Asp Gly Gly Xaa Gly Glu Leu Lys His Met Val Met Ser
-50 -45 -40

Phe Arg Val Ser Glu Leu Gln Val Leu Leu Gly Phe Ala Gly Arg Asn
-35
-30
-25

Lys Ser Gly Arg Lys His Glu Leu Leu Ala Lys Ala Leu His Leu Leu

-20 -15 -10 -5

Lys Ser Ser Cys Ala Pro Ser Val Gln Met Lys Ile Lys Glu Leu Tyr
1 5 10

Arg Arg Phe Pro Arg Lys Thr Leu Gly Pro Ser Asp Leu Ser Ser
15 20 25

Gly

- (2) INFORMATION FOR SEQ ID NO: 349:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 47 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -29..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.7 seq LCYLSIFCLGVLF/II
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 349:

Met Pro Cys Ile Ser Leu Leu Gly Leu Leu Tyr Asn Phe Val Gln Val -25 -20 -15

Leu Cys Tyr Leu Ser Ile Phe Cys Leu Gly Val Leu Phe Ile Ile Glu
-10 -5

Arg Gly Ser Leu Lys Val Ser Lys Leu Ile Cys Arg Pro Pro Gly
5 10 15

- (2) INFORMATION FOR SEQ ID NO: 350:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 47 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide

(B) LOCATION: -15..-1

- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 4.7 seq IAVLFCFFLLIIF/QT

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 350:

Met Lys Ile Ala Val Leu Phe Cys Phe Phe Leu Leu Ile Ile Phe Gln
-15 -5 1

Thr Asp Phe Gly Lys Asn Glu Glu Ile Pro Arg Lys Gln Arg Arg Lys

5 10 15

Ile Tyr His Arg Arg Leu Arg Lys Ser Ser Thr Ser His Lys Gln
20 25 30

- (2) INFORMATION FOR SEQ ID NO: 351:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 48 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -43..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.7

seg STWSSASLRGSWQ/QG

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 351:

Met Ala Lys Gln Lys Pro His Val Leu Gly Ser Arg Val Met Pro Ala
-40 -35 -30

Ser Cys Val Ser Glu Arg Arg Lys Pro Ser Phe Gln Val Ser Thr

Trp Ser Ser Ala Ser Leu Arg Gly Ser Trp Gln Gln Gly Met Pro Gly
-10 -5 1 5

- (2) INFORMATION FOR SEQ ID NO: 352:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 17 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo Sapiens
- (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig peptide
  - (B) LOCATION: -15..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.6

seq FLYLKSVFDVSLG/AR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 352:

Met Gly Phe Leu Tyr Leu Lys Ser Val Phe Asp Val Ser Leu Gly Ala -15 -5 1

Arg

- (2) INFORMATION FOR SEQ ID NO: 353:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 69 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -61..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.6

seq LLLLHGGGHSALS/WA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 353:

Met Arg Met Gly Pro Gly Arg Lys Arg Asp Phe Ser Pro Val Pro Trp
-60 -55 -50

Ser Gln Tyr Phe Glu Ser Met Glu Asp Val Glu Val Glu Asn Glu Thr -45 -35 -30

Gly Lys Asp Thr Phe Arg Val Tyr Lys Ser Gly Ser Glu Gly Pro Val -25 -20 -15

Leu Leu Leu His Gly Gly Gly His Ser Ala Leu Ser Trp Ala Val

Phe Thr Ala Ala Xaa 5 (2) INFORMATION FOR SEQ ID NO: 354:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 23 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -13..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.6

seq MIFLLYLLPSSEE/RR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 354:

Met Ile Phe Leu Leu Tyr Leu Leu Pro Ser Ser Glu Glu Arg Arg Lys
-10 -5 1

Leu Leu Phe Ser Pro His Arg
5 10

- (2) INFORMATION FOR SEQ ID NO: 355:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 70 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -61..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.6

seq LLLLHGGGHSALS/WA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 355:

Met Arg Met Gly Pro Gly Arg Lys Arg Asp Phe Ser Pro Val Pro Trp
-60 -55 - -50

Ser Gln Tyr Phe Glu Ser Met Glu Asp Val Glu Val Glu Asn Glu Thr
-45 -35 -30

Gly Lys Asp Thr Phe Arg Val Tyr Lys Ser Gly Ser Glu Gly Pro Val

Leu Leu Leu His Gly Gly Gly His Ser Ala Leu Ser Trp Ala Val

Phe Thr Ala Ala Thr Trp 5

- (2) INFORMATION FOR SEQ ID NO: 356:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 30 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -16..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.5

seq LLNLISILASIPS/QF

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 356:

Met Leu Ser Leu Leu Asn Leu Ile Ser Ile Leu Ala Ser Ile Pro Ser
-15 -5

Gln Phe Lys Pro Gln Phe Ser Lys Leu Pro Leu Ser Gly Arg
1 5

- (2) INFORMATION FOR SEQ ID NO: 357:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 28 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -25..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.5

seq LMLLWPVHPLLVG/HR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 357:

Met Gly Thr Thr Ser Asn Met Val Thr Thr Ile His Leu Met Leu Leu -25 -15 -10

Trp Pro Val His Pro Leu Leu Val Gly His Arg Gly
-5

- (2) INFORMATION FOR SEQ ID NO: 358:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 147 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION:  $-10\overline{1}..-1$
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.4 seq ISHILAFFAASDG/IV
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 358:

Met Gly Asp Pro Glu Arg Pro Glu Ala Ala Gly Leu Asp Gln Asp Glu
-100 -95 -90

Arg Ser Ser Ser Asp Thr Asn Glu Ser Glu Ile Lys Ser Asn Glu Glu
-85 -80 -75 -70

Pro Leu Leu Arg Lys Ser Ser Arg Arg Phe Val Ile Phe Pro Ile Gln
-65 -60 -55

Tyr Pro Asp Ile Trp Lys Met Tyr Lys Gln Ala Gln Ala Ser Phe Trp
-50
-45
-40

Thr Ala Glu Glu Val Asp Leu Ser Lys Asp Leu Pro His Trp Asn Lys
-35 -30 -25

Leu Lys Ala Asp Glu Lys Tyr Phe Ile Ser His Ile Leu Ala Phe Phe
-20 -15 -10

Ala Ala Ser Asp Gly Ile Val Asn Glu Asn Leu Val Glu Arg Phe Ser -5 1 5 10

Gin Glu Val Gln Val Pro Glu Ala Arg Cys Phe Tyr Gly Phe Gln Ile 15 20 25

Leu Ile Glu Asn Val His Ser Glu Met Tyr Ser Leu Leu Ile Asp Thr  $30 \hspace{1cm} 35 \hspace{1cm} 40$ 

Tyr Ile Arg 45

- (2) INFORMATION FOR SEQ ID NO: 359:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 65 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -16..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.4

seq GLFSLLPHPPCVG/RV

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 359:
- Met Asp Ala Gly Leu Phe Ser Leu Leu Pro His Pro Pro Cys Val Gly
  -15 -5
- Arg Val Leu Pro Gln Ser Arg Tyr His Leu His Pro Arg Ser Pro Leu
  1 5 10 15
- Val Glu Asp Thr Cys Phe Phe Gln Arg Leu Lys Lys Ile Leu Asn Lys 20 25 30
- Ile Gly Asn Leu Phe His Ser Thr Lys Ser Leu Cys Val Ser Leu Ala 35 40 45

Pro

- (2) INFORMATION FOR SEQ ID NO: 360:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 19 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -14..-1

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(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.4

seq LITLTYLIQGESA/RT

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 360:

Met Leu Ile Thr Leu Thr Tyr Leu Ile Gln Gly Glu Ser Ala Arg Thr

Thr Phe Glu

- (2) INFORMATION FOR SEQ ID NO: 361:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 31 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -26..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.4

seq RVQCLCAIPFAFS/LT

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 361:

Met Tyr Thr Gly Phe Arg Ile Glu Ala Thr Leu Leu Thr Arg Val Gln
-25 -15

Cys Leu Cys Ala Ile Pro Phe Ala Phe Ser Leu Thr Gly Ile Arg -10 -5 1 5

- (2) INFORMATION FOR SEQ ID NO: 362:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 56 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide

(B) LOCATION: -47..-1

- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 4.4 seq ISHILAFFAASDG/IV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 362:

Met Tyr Lys Gln Ala Gln Ala Ser Phe Trp Thr Ala Glu Glu Val Asp
-45
-40
-35

Leu Ser Lys Asp Leu Pro His Trp Asn Lys Leu Lys Ala Asp Glu Lys
-30
-25
-20

Tyr Phe Ile Ser His Ile Leu Ala Phe Phe Ala Ala Ser Asp Gly Ile
-15 -5 1

Val Asn Glu Asn Leu Val Glu Arg

- (2) INFORMATION FOR SEQ ID NO: 363:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 48 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -28..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.3

seq FLGLAAMASPSRN/SQ

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 363:

Met Leu Leu His Leu Cys Ser Val Lys Asn Leu Tyr Gln Asn Arg Phe -25 -20 -15

Leu Gly Leu Ala Ala Met Ala Ser Pro Ser Arg Asn Ser Gln Ser Arg
-10 -5 1

Arg Arg Cys Lys Glu Pro Leu Arg Tyr Ser Tyr Asn Pro Asp Gln Gly 5 10 15 20

- (2) INFORMATION FOR SEQ ID NO: 364:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 55 amino acids
    - (B) TYPE: AMINO ACID

- (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -18..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.3

seq WTCLKSFPSPTSS/HA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 364:

Met Pro Cys Pro Thr Trp Thr Cys Leu Lys Ser Phe Pro Ser Pro Thr
-15 -10 -5

Ser Ser His Ala Ser Ser Leu His Leu Pro Pro Ser Cys Thr Arg Leu  $1 \hspace{1.5cm} 5 \hspace{1.5cm} 10$ 

Thr Leu Thr Gln Thr Leu Arg Thr Gly Met His Leu Ser Arg Ala Leu 15 20 25 30

Gln Gly Thr Leu Thr Arg Gln

- (2) INFORMATION FOR SEQ ID NO: 365:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 73 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -33..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.3 seq LLGWGLNLTLGQG/AP
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 365:

Met Glu Asp Leu Phe Ser Pro Ser Ile Xaa Pro Pro Ala Pro Asn Ile
-30 -25 -20

Ser Val Pro Ile Leu Leu Gly Trp Gly Leu Asn Leu Thr Leu Gly Gln
-15 -10 -5

Gly Ala Pro Ala Ser Gly Pro Pro Ser Arg Arg Val Arg Leu Val Phe 1 5 10 15

Leu Gly Val Ile Leu Val Val Ala Val Ala Xaa Asn Thr Thr Val Leu 20 25 30

Cys Arg Leu Cys Gly Gly Gly Pro 35 40

- (2) INFORMATION FOR SEQ ID NO: 366:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 55 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig peptide
    - (B) LOCATION: -52..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.1

seq VMLETCGLLVSLG/HP

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 366:
- Met Ala Glu Thr Lys Asp Ala Ala Gln Met Leu Val Thr Phe Lys Asp
  -50 -45 -40
- Val Ala Val Thr Phe Thr Arg Glu Glu Trp Arg Gln Leu Asp Leu Ala
  -35
  -30
  -25
- Gln Arg Thr Leu Tyr Arg Glu Val Met Leu Glu Thr Cys Gly Leu Leu
  -20 -15 -10 -5

Val Ser Leu Gly His Pro Arg

(2) INFORMATION FOR SEQ ID NO: 367:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 45 amino acids
- (B) TYPE: AMINO ACID
- (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain

(ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: -13..-1
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 4.1

seq MLILSQNIAQLEA/QV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 367:

Met Leu Ile Leu Ser Gln Asn Ile Ala Gln Leu Glu Ala Gln Val Glu
-10 -5 1

Lys Val Thr Lys Glu Lys Ile Ser Ala Ile Asn Gln Leu Glu Glu Asn
5 10 15

Ser Lys Pro Ala Gly Phe Ser Gly Lys Trp Met Ser Gln 20 25 30

- (2) INFORMATION FOR SEQ ID NO: 368:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 97 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -21..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.1

seq GLWAHSWTCSCSA/AX

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 368:

Met Leu Cly Ala Ser Ala Gln Gly Leu Trp Ala His Ser Trp Thr

Cys Ser Cys Ser Ala Ala Xaa Arg Ser Val His Pro Gly Gly Asp Trp
-5 1 10

Met Gln Gln Phe Gln Ala Gly Phe Leu Pro Pro Gln Val Pro Ala His
15 20 25

Leu Ser Leu Thr Trp Asp Val Ser Leu Leu Pro Pro Cys Leu Val Pro 30 35 40

Lys Ala Leu Glu Phe Val Val His Phe Leu Lys Asn Asp Ile Phe Tyr 45 50 55

Leu Thr Gln Tyr Ile Lys Asn Val Ile Ser Glu Cys Thr Phe Ser Phe 60 65 70 75

Phe

(2) INFORMATION FOR SEQ ID NO: 369:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 68 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -15..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.1

seq APLELSCWGGGWG/LP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 369:

Met Ala Ala Pro Leu Glu Leu Ser Cys Trp Gly Gly Gly Trp Gly Leu
-15 -5 1

Pro Ser Val His Ser Glu Ser Leu Val Val Met Ala Tyr Ala Lys Phe 5 10

Ser Gly Ala Pro Leu Lys Val Asn Val Ile Asp Asn Thr Trp Arg Gly 20 25 30

Ser Arg Gly Asp Val Pro Ile Leu Thr Thr Glu Asp Asp Met Val Ser 35 40 45

Gln Pro Ala Arg 50

- (2) INFORMATION FOR SEQ ID NO: 370:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 69 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (E) LOCATION: -20..-1

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.1

seq LGFLNCYIAVARS/GG

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 370:

Met Ser Xaa Val Gly Ile Asp Leu Gly Phe Leu Asn Cys Tyr Ile Ala -20 -15 -10 -5

Val Ala Arg Ser Gly Gly Ile Glu Thr Ile Ala Asn Glu Tyr Ser Asp 1 5 10

Arg Cys Thr Pro Ala Cys Ile Ser Leu Gly Ser Arg Thr Arg Ala Ile 15 20 25

Gly Asn Ala Ala Lys Ser Gln Ile Val Thr Asn Val Arg Asn Thr Ile  $30 \hspace{1cm} 35 \hspace{1cm} 40$ 

His Gly Phe Lys Lys

- (2) INFORMATION FOR SEQ ID NO: 371:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 43 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -19..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.1

seq FVVFSTMFTASSP/GE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 371:

Met Glu Tyr Ser Lys Xaa Phe Val Val Phe Ser Thr Met Phe Thr Ala-15 -10 -5

Ser Ser Pro Gly Glu Asp Phe Pro Pro Phe Phe Ser Gln Met Xaa Arg

Leu Ser Arg Asn Tyr Phe Pro Cys Pro Pro Xaa 15 20

- (2) INFORMATION FOR SEQ ID NO: 372:
  - (i) SEQUENCE CHARACTERISTICS:

PCT/IB98/01236 WO 99/06552

(A) LENGTH: 72 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig peptide
  - (B) LOCATION: -40..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.1

seq LPFRLPWASTATA/RC

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 372:

Met Pro Met Ala Ser Ser Pro Pro Pro Ser Pro His Pro Gln Glu Pro

Ala Pro Leu Leu Pro Ser Leu Pro Arg Leu Ser Leu Pro Phe Arg Leu

Pro Trp Ala Ser Thr Ala Thr Ala Arg Cys Pro Pro Ser Pro Leu Gly

Ser Leu Xaa Leu Met Leu Cys Ile Pro Thr Gly Phe Thr Pro Thr Gln

Pro Arg Ala Pro Arg Pro Pro Gly 30

- (2) INFORMATION FOR SEQ ID NO: 373:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 55 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -38..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4

seq WALGLKFLSSSSQ/NF

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 373:

Met Gln His Val Xaa Gly His Xaa Pro Asp Pro Ile Ala Ile Met Tyr

-35 -30 -25

Val Cys Pro Pro Cys Gly His Thr Thr Trp Ala Leu Gly Leu Lys Phe
-20 -15 -10

Leu Ser Ser Ser Ser Gln Asn Phe Cys Ala Pro Val Leu Phe Leu Ile
-5 1 5 10

Leu His Thr Gly Gly Gln Arg

- (2) INFORMATION FOR SEQ ID NO: 374:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 53 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -30..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4

seq AGFLKCLLLSSLQ/SY

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 374:

Met Gly Trp Glu Met Thr Cys Ile Lys Ser Phe Phe Trp Ala Arg Ser -30 -25 -20 -15

His Ala Gly Phe Leu Lys Cys Leu Leu Leu Ser Ser Leu Gln Ser Tyr
-10 -5 1

Lys Glu Ala Ala Val Ile Phe Pro Leu Thr Asp Leu Lys Leu Lys
5 10 15

Asp Tyr Gly Glu Trp 20

- (2) INFORMATION FOR SEQ ID NO: 375:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 38 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -36..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4

seq VQLSFAATTPVLA/DK

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 375:

Met Val Phe Gly Gly Val Cys Pro Ser Val Thr Ser Ile Ile Ala Glu
-35 -25

Ser Leu Gln Gly Trp Asn Leu Val Gln Leu Ser Phe Ala Ala Thr Thr
-20 -15 -10 -5

Pro Val Leu Ala Asp Lys

- (2) INFORMATION FOR SEQ ID NO: 376:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 24 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -16..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4

seq ITWSLLFLYQCSL/HF

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 376:

Met His Phe Ile Thr Trp Ser Leu Leu Phe Leu Tyr Gln Cys Ser Leu -15 -5

His Phe Ile Ile Ile Lys Ala Gly
1 5

- (2) INFORMATION FOR SEQ ID NO: 377:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 41 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -36..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.9

seq CWPSVASPSSSWS/SP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 377:

Met Ser Gly Ala Ser Pro Ile Glu Arg Thr Pro Met Glu Glu Ala Pro -35 -25

Ser Ser Cys Pro Thr Ser Ser Cys Trp Pro Ser Val Ala Ser Pro Ser -20 -15 -10 -5

Ser Ser Trp Ser Ser Pro Trp Ala Ser

- (2) INFORMATION FOR SEQ ID NO: 378:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 50 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -37..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.9

seq PGPSLRLFSGSQA/SV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 378:

Met Glu Trp Ala Gly Lys Gln Arg Asp Phe Gln Val Arg Ala Ala Pro
-35 -30 -25

Gly Trp Asp His Leu Ala Ser Phe Pro Gly Pro Ser Leu Arg Leu Phe
-20 -15 -10

Ser Gly Ser Gln Ala Ser Val Cys Ser Leu Cys Ser Gly Phe Gly Ala -5 5 10

Gln Glu

(2) INFORMATION FOR SEQ ID NO: 379:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 67 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: -60..-1

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 3.9

seq AKVVSLSLQTSSA/HH

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 379:

Met Ile Ala Phe Phe Asp Glu Asp Asn Pro Arg Lys Arg Arg Ser Tyr
-60 -55 -50 -50

Ser Phe Thr Gln Ser Ala Gly Ile Leu Cys Gln Glu Thr Thr Tyr Ser -40 -35 -30

Thr Pro His Thr Lys Leu Glu Lys Ala Lys Ser Pro Thr Ala Asp Ala
-25
-20
-15

Lys Val Val Ser Leu Ser Leu Gln Thr Ser Ser Ala His His Arg Gly
-10 -5 1

Gly Xaa Gly

- (2) INFORMATION FOR SEQ ID NO: 380:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 87 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (7) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -48..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.9

seq ALFCTLPCPVERG/QQ

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 380:

Met Gly Lys Ser Ile Xaa Ser Leu Cys Ser Val Xaa Leu Lys Ala Arg

Leu Lys Gly Xaa Leu Glu Ala Val His Leu Cys Leu Arg Ala Gln Lys
-30 -25 -20

Arg Arg Thr Ala Leu Phe Cys Thr Leu Pro Cys Pro Val Glu Arg Gly
-15 -5

Gln Gln Val Pro Gly Xaa Xaa Xaa Arg Leu Arg Leu Ala Ser Pro Ser 1 5 10 15

Val Ala Lys Val Phe Gln Cys Phe Leu Ser Lys Leu Cys Val Trp Asn 20 25 30

Ile Lys Asp Gly Leu Ser Arg 35

- (2) INFORMATION FOR SEQ ID NO: 381:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 53 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -15..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.9 seq LHMTLFRVPFTFS/XF
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 381:

Met Cys Leu His Met Thr Leu Phe Arg Val Pro Phe Thr Phe Ser Xaa -15 -5 1

Phe Trp Lys Gly Ala Gly Arg Gln Glu Glu Cys Ser Phe Lys Pro Ser 5 10 15

Leu Tyr Tyr Lys Leu Ile Met Val Leu Lys Ile Ala Leu Leu Leu 20 25 30

Ser Pro Pro Pro Lys

(2) INFORMATION FOR SEQ ID NO: 382:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 23 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -14..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.9

seq LNILKTLTSAALP/SP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 382:

Met Leu Asn Ile Leu Lys Thr Leu Thr Ser Ala Ala Leu Pro Ser Pro
-10 -5 1

Ser Pro Arg Pro Asn Lys Arg

- (2) INFORMATION FOR SEQ ID NO: 383:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 51 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -40..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.9

seq SPLLCLYHPPVYT/ST

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 383:

Met Arg Ala Arg Val Trp Pro Arg Ser His Gly Ile Pro Val Pro Ser
-40 -35 -30 -25

Phe Leu Ser Lys Ser Ser Leu Ser His Thr Pro Ser Pro Leu Leu Cys
-20
-15
-10

Let Tyr His Pro Pro Val Tyr Thr Ser Thr Thr Thr Pro Ser Ile Pro

**-**5 1 5

Pro Arg Leu 10

(2) INFORMATION FOR SEQ ID NO: 384:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 51 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -36..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.9

seq SLCLSLLIPGPKP/LV

....

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 384:

Met Trp Asn Ala Val Ala Ile Ile Cys Asn Gly Ser Trp Cys Gln Thr
-35
-30
-25

Xaa Ser Thr Ser Gly Leu Glu Ser Leu Cys Leu Ser Leu Leu Ile Pro
-20 -15 -10 -5

Gly Pro Lys Pro Leu Val Ser Val Gly Ile Asn Gln Leu Leu Thr
1 5 10

Ser Ser Arg 15

- (2) INFORMATION FOR SEQ ID NO: 385:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 23 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -14..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 3.9

seq LRLGLFKISWARC/LS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 385:

Met Leu Arg Leu Gly Leu Phe Lys Ile Ser Trp Ala Arg Cys Leu Ser

Tyr Ser Lys Thr Gln Xaa Glu

- (2) INFORMATION FOR SEQ ID NO: 386:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 61 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -36..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.9

seq VVEILPYLPCLTA/RD

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 386:

Met Pro Phe Ala Glu Asp Lys Thr Tyr Lys Tyr Ile Cys Arg Asn Phe
-35
-30
-25

Ser Asn Phe Cys Asn Val Asp Val Val Glu Ile Leu Pro Tyr Leu Pro -20 -15 -10 -5

Cys Leu Thr Ala Arg Asp Gln Asp Arg Leu Arg Ala Thr Cys Thr Leu

Ser Gly Asn Arg Asp Thr Leu Trp His Leu Phe Asn Thr
15 20 25

- (2) INFORMATION FOR SEQ ID NO: 387:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 40 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:

- · (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -36..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.8

seq GTDSLSFLPPCPC/CP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 387:

Met Pro Gly Ser Ser Gly Leu Arg Phe Ile Cys Lys Ser Arg Asn His -35 -30 -25

Pro Gln Phe Gly Ser Phe Ser Gly Thr Asp Ser Leu Ser Phe Leu Pro
-20 -15 -10 -5

Pro Cys Pro Cys Cys Pro Ala Ala

- (2) INFORMATION FOR SEQ ID NO: 388:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 66 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -57..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.8

seq QLXLVMEFCGAGS/VT

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 388:

Met Asp Val Thr Gly Asp Glu Glu Glu Glu Ile Lys Gln Glu Ile Asn
-55 -50 -45

Met Leu Lys Lys Tyr Ser His His Arg Asn Ile Ala Thr Tyr Tyr Gly
-40 -35 -30

Ala Phe Ile Lys Lys Asn Pro Pro Gly Met Asp Asp Gln Leu Xaa Leu
-25 -10 -15

Val Met Glu Phe Cys Gly Ala Gly Ser Val Thr Asp Leu Ile Lys Asn -5 5

Thr Gly

365

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(2) INFORMATION FOR SEQ ID NO: 389:
```

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 26 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -24..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.8

seg KLFLVFLLNICKG/IV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 389:

Met Ile Phe Gly Leu Tyr Phe Val Leu Ala Val Lys Leu Phe Leu Val
-20 -15 -10

Phe Leu Leu Asn Ile Cys Lys Gly Ile Val

- (2) INFORMATION FOR SEQ ID NO: 390:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 46 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -34..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.6

seq IKCSSWISSLASG/IP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 390:

Met Arg Lys Lys Arg Val Glu Glu Leu Ile Val Phe Pro Gly Glu Val
-30 -25 -20

Thr Ser Phe Ser Ser Ile Lys Cys Ser Ser Trp Ile Ser Ser Leu Ala
-15 -10 -5

Ser Gly Ile Pro His Ser Leu Gly Phe Ser Leu Pro Pro Gly
1 5 10

(2) INFORMATION FOR SEQ ID NO: 391:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 57 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -28..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.6

seq ACLFSXFLAVSRH/PN

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 391:

Met Pro Ser Ser Ser Leu Ala Glu Leu Cys Leu Met Gln Gln Asp Ala

Cys Leu Phe Ser Xaa Phe Leu Ala Val Ser Arg His Pro Asn Tyr Xaa -10 -5 1

Cys Ser Ile Ser Thr Lys Gly Glu Val Arg Glu Lys Leu Val Pro Trp 5 10 15 20

Ile Thr His Gln Met Ala Arg Met Leu 25

- (2) INFORMATION FOR SEQ ID NO: 392:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 46 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -40..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.6

seq LQMRMQLPCLVLG/EE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 392:

Met Asp Leu Trp Ser Cys Leu Phe Pro Val Met Leu Met Glu Pro Ser
-40 -35 -30 -25

Lys Gly Leu Glu Asp Ser Glu Trp Lys Met Ala Leu Gln Met Arg Met -20 -15 -10

Gln Leu Pro Cys Leu Val Leu Gly Glu Glu Gln Thr Leu Gly
-5 1 5

- (2) INFORMATION FOR SEQ ID NO: 393:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 37 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -27..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.6

seq AVPLPTTSTLTSA/ST

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 393:

Met Ser Gly Lys Gly Lys Cys Arg Pro Ile Ala Leu Arg Arg Ala Val -25 -20 -15

Pro Leu Pro Thr Thr Ser Thr Leu Thr Ser Ala Ser Thr Gly Phe Leu
-10 -5 1 5

Trp Ile Leu Lys Glu

- (2) INFORMATION FOR SEQ ID NO: 394:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 26 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens

· (F) TISSUE TYPE: Brain

#### (ix) FEATURE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: -18..-1
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 3.6

seq IQKSSGLFCPSQA/QS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 394:

Met Thr Pro Lys Ala Ile Gln Lys Ser Ser Gly Leu Phe Cys Pro Ser
-15 -10 -5

Gln Ala Gln Ser Ala Arg Pro Ala Glu Lys 1 5

#### (2) INFORMATION FOR SEQ ID NO: 395:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 103 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -72..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.6

seq CTSLLQLYDASNS/EW

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 395:

Met Pro Asp Gln Phe Asp Gln Ala Val Val Leu Asn Gln Leu Arg Tyr
-70 -65 -60

Ser Gly Met Leu Glu Thr Val Arg Ile Arg Lys Ala Gly Tyr Ala Val -55 -50 -45

Arg Arg Pro Phe Gln Asp Phe Tyr Lys Arg Tyr Lys Val Leu Met Arg -40 -35 -30 -25

Asn Leu Ala Leu Pro Glu Asp Val Arg Gly Lys Cys Thr Ser Leu Leu
-20 -15 -10

Gln Leu Tyr Asp Ala Ser Asn Ser Glu Trp Gln Leu Gly Lys Thr Lys
-5 1 5

Val Phe Leu Arg Glu Ser Leu Glu Gln Lys Leu Glu Lys Arg Arg Glu 10 15 20

Glu Glu Val Ser His Ala Gly

WO 99/06552

25 30

(2) INFORMATION FOR SEQ ID NO: 396:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 19 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: -15..-1

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 3.6

seq LVSFFLELNVLQQ/WP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 396:

Met Cys Leu Val Ser Phe Phe Leu Glu Leu Asn Val Leu Gln Gln Trp -10 -15 -5

Pro Ala Gly

(2) INFORMATION FOR SEQ ID NO: 397:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 44 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: -13..-1

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 3.6

seq MRSLACLTPCGHA/GS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 397:

Met Arg Ser Leu Ala Cys Leu Thr Pro Cys Gly His Ala Gly Ser Arg -10

Leu Gln Ser Ser Leu Ser Lys Tyr Leu Val Leu Pro Asn Leu Glu Cys
5 10 15

Leu Phe Phe Leu Phe Leu Ile Ser Asn Arg Arg Trp 20 25 30

- (2) INFORMATION FOR SEQ ID NO: 398:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 82 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -13..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.5

seq MHLLSNWANPASS/RR

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 398:
- Met His Leu Leu Ser Asn Trp Ala Asn Pro Ala Ser Ser Arg Arg Pro
  -10 -5 1
- Ser Met Ala Ala Ser Gly Thr Ser Trp Ile Ser Ser Thr Leu Ala His
- Ser Leu Ser Leu Arg Asp Val Ser Glu Arg Leu Cys Ser Cys Trp Arg 20 25 30 35
- Thr Ile Ser Met Gly Pro Cys Ala Arg Gly Ser Pro Met Asn Ser Ser 40 45 50
- Gly Val His Arg Lys Ser Ser Arg Leu Phe Tyr Ile Arg Thr Pro Met
  55 60 65

Arg Arg

- (2) INFORMATION FOR SEQ ID NO: 399:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 118 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -24..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.5

seq FAMLHSVWRLIPA/FR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 399:

Met Trp Ser Gly Lys Trp Ala Leu Val Ser Pro Phe Ala Met Leu His

Ser Val Trp Arg Leu Ile Pro Ala Phe Arg Gly Tyr Ala Gln Gln Asp
-5 1 5

Ala Gln Glu Phe Leu Cys Glu Leu Leu Asp Lys Ile Gln Arg Glu Leu 10 20

Glu Thr Thr Gly Thr Ser Leu Pro Ala Leu Ile Pro Thr Ser Gln Arg 25 30 35 40

Lys Leu Ile Lys Gln Val Leu Asn Val Val Asn Asn Ile Phe His Gly 45 50 55

Gln Leu Leu Ser Gln Val Thr Cys Leu Ala Cys Asp Asn Lys Ser Asn 60 65 70

Thr Ile Glu Pro Phe Trp Asp Leu Ser Leu Glu Xaa Pro Glu Arg Tyr
75 80 85

Gln Cys Ser Xaa Lys Gly 90

- (2) INFORMATION FOR SEQ ID NO: 400:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 41 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -20..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.5
      - seq KFCLICLLTFIFH/HC
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 400:

Met Lys Val His Met His Thr Lys Phe Cys Leu Ile Cys Leu Leu Thr -20 -15 -10 -5

Phe Ile Phe His His Cys Asn His Cys His Glu Glu His Asp His Gly
1 5 10

Pro Glu Ala Leu His Arg Gln Gln Gly 15 20

- (2) INFORMATION FOR SEQ ID NO: 401:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 49 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -24..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.5

seq ALSLFYTADTSHG/SE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 401:

Met Gly Arg Arg His Trp Val Leu Thr His Ser Ala Leu Ser Leu Phe -20 -15 -10

Tyr Thr Ala Asp Thr Ser His Gly Ser Glu Lys Pro Tyr Leu Ser Leu
-5 5

Phe Gly Arg Glu Gly Gly Arg Glu Gly Ser Asn Pro Lys Tyr Tyr Ser 10 15 20

Phe 25

- (2) INFORMATION FOR SEQ ID NO: 402:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 93 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -16..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 12.5

seq FVVLLALVAGVLG/NE

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 402:
- Met Ala Val Phe Val Val Leu Leu Ala Leu Val Ala Gly Val Leu Gly
  -15
  -5
- Asn Glu Phe Ser Ile Leu Lys Ser Pro Gly Ser Val Val Phe Arg Asn 1 5 10 15
- Gly Asn Trp Pro Ile Pro Gly Glu Arg Ile Pro Asp Val Ala Ala Leu 20 25 30
- Ser Met Gly Phe Ser Val Lys Glu Asp Leu Ser Trp Pro Gly Leu Ala 35 40 45
- Val Gly Asn Leu Phe His Arg Pro Arg Ala Ser Val Met Val Met Val 50 55 60
- Lys Gly Val Asn Asn Xaa Pro Leu Pro Pro Xaa Trp Xaa 65 70 75
- (2) INFORMATION FOR SEQ ID NO: 403:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 65 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -16..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 10.1

seq LLLQLAVLGAALA/AA

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 403:
- Met Ala Pro Leu Leu Gln Leu Ala Val Leu Gly Ala Ala Leu Ala -15 -5
- Ala Ala Ala Leu Val Leu Ile Ser Ile Val Ala Phe Thr Thr Ala Thr 1 5 10 15
- Lys Met Pro Ala Leu His Arg His Glu Glu Glu Lys Phe Phe Leu Asn 20 25 30

Ala Lys Gly Gln Lys Glu Thr Leu Pro Ser Ile Trp Asp Ser Pro Thr
35 40 45

Arg

- (2) INFORMATION FOR SEQ ID NO: 404:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 84 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -50..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 9.8

seq LLRLLQLVSTCVA/FS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 404:

Met Pro Val Thr Val Thr Arg Thr Thr Ile Thr Thr Thr Thr Ser
-50 -45 -40 -35

Ser Ser Gly Leu Gly Ser Pro Met Ile Val Gly Ser Pro Arg Ala Leu
-30 -25 -20

Thr Gln Pro Leu Gly Leu Leu Arg Leu Leu Gln Leu Val Ser Thr Cys
-15
-10
-5

Val Ala Phe Ser Leu Val Ala Ser Val Gly Ala Trp Thr Giy Ser Met

1 5 10

Gly Asn Trp Ser Met Phe Thr Trp Cys Phe Cys Phe Ser Val Thr Leu 15 20 25 30

Ile Ile Leu Ile

- (2) INFORMATION FOR SEQ ID NO: 405:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 72 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -18..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 9.2

seq VFLCSLLAPMVLA/SA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 405:

Met Glu Leu Val Leu Val Phe Leu Cys Ser Leu Leu Ala Pro Met Val -15 -10 -5

Leu Ala Ser Ala Ala Glu Lys Glu Lys Glu Met Asp Pro Phe His Tyr  $1 \hspace{1cm} 5 \hspace{1cm} 10$ 

Asp Tyr Gln Thr Leu Arg Ile Gly Gly Leu Val Phe Ala Val Val Leu 15 20 25 30

Phe Ser Val Gly Ile Leu Leu Ile Leu Ser Arg Arg Cys Lys Cys Ser 35 40 45

Phe Asn Gln Lys Pro Arg Asn Arg

- (2) INFORMATION FOR SEQ ID NO: 406:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 70 amino acids
      - (B) TYPE: AMINO ACID
      - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -46..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7.9

seq LLGLLSAEQLAEA/SV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 406:

Met Gly Pro Ile Trp Ser Ser Tyr Tyr Gly Asn Cys Arg Ser Leu Leu
-45 -40 -35

Phe Val Met Asp Ala Ser Asp Pro Thr Gln Leu Ser Ala Ser Cys Val -30 -25 -20 -15

Gin Leu Leu Gly Leu Leu Ser Ala Glu Gln Leu Ala Glu Aia Ser Val

Leu Ile Leu Phe Asn Lys Ile Asp Leu Pro Cys Tyr Met Ser Thr Glu 10

Glu Met Lys Ser Leu Ile 20

- (2) INFORMATION FOR SEQ ID NO: 407:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 57 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -47..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7.9

seq LLLPRVLLTMASG/SP

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 407:
- Met Ser Gly Gly Arg Ala Pro Ala Val Leu Leu Gly Gly Val Ala Ser -45 -40
- Leu Leu Ser Phe Val Trp Met Pro Ala Leu Leu Pro Val Ala Ser -25
- Arg Leu Leu Leu Pro Arg Val Leu Leu Thr Met Ala Ser Gly Ser -15 -10 -5

Pro Pro Thr Gln Pro Ser Pro Ala Trp 5

- (2) INFORMATION FOR SEQ ID NO: 408:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 92 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -40..-1

WO 99/06552 PCT/IB98/01236

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 7.9 seq SLLLLFGGQFASS/QE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 408:

Met Ala Leu Ser Cys Thr Leu Asn Arg Tyr Leu Leu Leu Met Ala Gln
-40
-35
-30
-25

Glu His Leu Glu Phe Arg Leu Pro Glu Ile Xaa Ser Leu Leu Leu Leu -20 -15 -10

Phe Gly Gly Gln Phe Ala Ser Ser Gln Glu Thr Tyr Gly Lys Ser Pro
-5 1 5

Phe Trp Ile Leu Ser Ile Pro Ser Glu Asp Ile Ala Arg Asn Leu Met
10 15 20

Lys Arg Thr Val Cys Ala Lys Ser Ile Phe Glu Leu Trp Gly His Gly 25 30 35 40

Gln Ser Pro Glu Glu Leu Tyr Ser Ser Leu Lys Asn
45

- (2) INFORMATION FOR SEQ ID NO: 409:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 91 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -53..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7.1

seq IAVGLGVAALAFA/GR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 409:

Met Ala Ala Arg Gly Val Ile Ala Pro Val Gly Glu Ser Leu Arg Tyr
-50 -45 -40

Ala Glu Tyr Leu Gln Pro Ser Ala Lys Arg Pro Asp Ala Asp Val Asp
-35
-30
-25

Gln Gln Arg Leu Val Arg Ser Leu Ile Ala Val Gly Leu Gly Val Ala
-20 -15 -10

Ala Leu Ala Phe Ala Gly Arg Tyr Ala Phe Arg Ile Trp Lys Pro Leu
-5 1 10

Glu Gln Val Ile Thr Glu Thr Ala Lys Lys Ile Ser Thr Pro Ser Phe
15 20 25

Ser Ser Tyr Tyr Lys Gly Gly Phe Glu Arg Arg 30 35

- (2) INFORMATION FOR SEQ ID NO: 410:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 31 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -28..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 6.6

seg VLGXLFLGGLCRG/WD

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 410:

Met Arg Met Cys Ala Gly Ser Ile Tyr Lys Ser Ala Thr Gln Ala Val -25 -20 -15

Leu Gly Xaa Leu Phe Leu Gly Gly Leu Cys Arg Gly Trp Asp Ala
-10 -5 1

- (2) INFORMATION FOR SEQ ID NO: 411:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 80 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (2) LOCATION: -73..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 6.3

seq TLIMLLSWQLSVS/SV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 411:

Met Ala Glu Arg Arg Pro Leu Ser Pro Ile Pro Ser Xaa Arg Arg
-75 -70 -65

Pro Ser Glu Pro Ser Arg Pro Arg Pro Ala Ala Gly Xaa Arg Ser
-60 -55 -50

Leu Pro Arg Pro Gly Asp Glu Glu Leu Gln Leu Pro Cys Ala Val His
-45 -40 -35

Asp Leu Ile Phe Trp Arg Asp Val Lys Lys Thr Gly Phe Val Phe Gly
-30 -25 -20 -15

Thr Thr Leu Ile Met Leu Leu Ser Trp Gln Leu Ser Val Ser Ser Val -10 -5 1

## (2) INFORMATION FOR SEQ ID NO: 412:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 133 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig peptide
  - (B) LOCATION: -109..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 6

seq LQLLLGMTASAVA/AL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 412:

Met Ala Ala Pro Val Leu Leu Arg Val Ser Val Pro Arg Trp Glu Arg
-105 -100 -95

Val Ala Arg Tyr Ala Val Cys Ala Ala Gly Ile Leu Leu Ser Ile Tyr
-90 -85 -80

Ala Tyr His Val Glu Arg Glu Lys Glu Arg Asp Pro Glu His Arg Ala
-75 -70 -65

Leu Cys Asp Leu Gly Pro Trp Val Lys Cys Ser Ala Ala Leu Ala Ser
-60 -55 -50

Arg Trp Gly Arg Gly Phe Gly Leu Leu Gly Ser Ile Phe Gly Lys Asp
-45 -35 -30

Gly Val Leu Asn Gln Pro Asn Ser Val Phe Gly Leu Ile Phe Tyr Ile
-25
-20
-15

Leu Gln Leu Leu Gly Met Thr Ala Ser Ala Val Ala Ala Leu Ile

. -10 -5 1

Leu Met Thr Ser Ser Ile Met Ser Val Val Gly Ser Cys Thr Trp Pro
5 10 15

Thr Phe Cys Thr Thr 20

- (2) INFORMATION FOR SEQ ID NO: 413:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 106 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -34..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.9 seq LGAAALALLLANT/DV
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 413:

Met Ser Phe Leu Gln Asp Pro Ser Phe Phe Thr Met Gly Met Trp Ser -30 -25 -20

Ile Gly Ala Gly Ala Leu Gly Ala Ala Ala Leu Ala Leu Leu Leu Ala
-15
-10
-5

Asn Thr Asp Val Phe Leu Ser Lys Pro Xaa Lys Ala Ala Leu Glu Tyr  $1 \hspace{1cm} 5 \hspace{1cm} 10$ 

Leu Glu Asp Ile Asp Leu Lys Thr Leu Glu Lys Glu Pro Arg Thr Phe
15 20 25 30

Lys Ala Lys Glu Leu Trp Glu Lys Asn Gly Ala Val Ile Met Ala Val 35 40 45

Arg Arg Pro Giy Cys Phe Leu Cys Arg Glu Glu Ala Ala Asp Leu Ser 50 55 60

Ser Leu Lys Ser Met Leu Asp Gln Leu Gly 65 70

- (2) INFORMATION FOR SEQ ID NO: 414:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 56 amino acids
    - (B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: -37..-1

- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 4.9

seq MLIMLGIFFNVHS/AV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 414:

Met Ala Ser Leu Leu Cys Cys Gly Pro Lys Leu Ala Ala Cys Gly Ile
-35
-30
-25

Val Leu Ser Ala Trp Gly Val Ile Met Leu Ile Met Leu Gly Ile Phe
-20 -15 -10

Phe Asn Val His Ser Ala Val Leu Ile Glu Asp Val Pro Phe Thr Glu -5 10

Lys Asp Phe Glu Asn Gly Pro Arg 15

- (2) INFORMATION FOR SEQ ID NO: 415:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 32 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -20..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.9

seq XSLFLHAVSSSFT/QL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 415:

Met Ile Leu Pro Tyr Arg Met Xaa Ser Leu Phe Leu His Ala Val Ser -20 -15 -10 -5

Ser Ser Phe Thr Gln Leu Arg Ser Cys Gln Gly Asp Arg Val Trp Arg
1 5 10

(2) INFORMATION FOR SEQ ID NO: 416:

(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 78 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -63..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.8

seq LYTVRALAGRAWA/AV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 416:

Met Ala Thr Leu Val Glu Leu Pro Asp Ser Val Leu Leu Glu Ile Phe

Ser Tyr Leu Pro Val Arg Asp Arg Ile Arg Ile Ser Arg Val Cys His

Arg Trp Lys Arg Leu Val Asp Asp Arg Trp Leu Trp Arg His Val Asp

Leu Thr Leu Tyr Thr Val Arg Ala Leu Ala Gly Arg Ala Trp Ala Ala

Val Ala Val Pro Gly Xaa Arg Arg Pro Pro Leu Pro Pro Trp 5 . 10

- (2) INFORMATION FOR SEQ ID NO: 417:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 92 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -41..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.7

seq LFSCFCFLSHKFG/KK

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 417:

Met Lys Asn Ala Cys Ile Val Leu Pro Pro Thr Pro Pro Pro Ser Leu
-40 -35 -30

Gln Pro Ser Ala Ser Leu Leu Ala Pro Asn Arg Phe Leu Phe Ser Cys
-25 -15 -10

Phe Cys Phe Leu Ser His Lys Phe Gly Lys Lys Val Ile Tyr Phe Asn
-5

Tyr Leu Ser Glu Leu His Glu His Leu Lys Tyr Asp Gln Leu Val Ile 10 15 20

Pro Pro Glu Val Leu Arg Tyr Asp Glu Lys Leu Gln Ser Leu His Glu 25 30

Gly Arg Thr Pro Xaa Pro Thr Lys Thr Pro Pro Gly 40 45 50

- (2) INFORMATION FOR SEQ ID NO: 418:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 55 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -28..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.6 seq PLQWSLLVAVVAG/SV
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 418:

Met Ala Phe Gly Leu Gln Met Phe Ile Gln Arg Lys Phe Pro Tyr Pro
-25 -20 -15

Leu Gln Trp Ser Leu Leu Val Ala Val Val Ala Gly Ser Val Val Ser -10 -5 1

Tyr Gly Val Thr Arg Val Xaa Ser Glu Lys Cys Asn Asn Leu Trp Leu 5 10 15 20

Phe Leu Glu Thr Gly Leu Gly

25

(2) INFORMATION FOR SEQ ID NO: 419:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 69 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: -53..-1

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.5

seq LLWTPLLSPGSLR/VI

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 419:

Met Tyr Cys Lys Ile Leu Val Leu Met Leu His Thr Glu Leu Ile Arg
-50 -45 -40

Thr Asp Tyr Ser Ser Val Asp Gln Leu Leu Leu Asn Tyr Pro Ala Glu
-35
-30
-25

Glu Gly Leu Gly Arg Glu Arg Ser Leu Leu Trp Thr Pro Leu Leu Ser
-20 -15 -10

Pro Gly Ser Leu Arg Val Ile Leu Glu Ser Arg Glu Val Pro Val Ser -5 1 5 10

Leu Trp Pro Gln Thr

- (2) INFORMATION FOR SEQ ID NO: 420:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 86 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -26..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.5

seq ENSLIILLQGLQG/RV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 420:

Met Ala Val Ser His Ser Val Lys Glu Arg Thr Ile Ser Glu Asn Ser -25 -20 -15

Leu Ile Ile Leu Leu Gln Gly Leu Gln Gly Arg Val Thr Thr Val Asp
-10 -5 1 5

Leu Arg Asp Glu Ser Val Ala His Gly Arg Ile Asp Xaa Val Asp Ala 10 15 20

Phe Met Asn Ile Arg Leu Ala Lys Val Thr Tyr Thr Asp Arg Trp Gly 25 30 35

His Gln Val Lys Leu Asp Asp Leu Phe Val Thr Gly Arg Asn Val Arg
40 45 50

Tyr Val His Ile Pro Asp

- (2) INFORMATION FOR SEQ ID NO: 421:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 91 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -23..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.4

seq QFILLGTTSVVTA/AL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 421:

Met Glu Ser Gly Gly Arg Pro Ser Leu Cys Gln Phe Ile Leu Leu Gly
-20 -15 -10

Thr Thr Ser Val Val Thr Ala Ala Leu Tyr Ser Val Tyr Arg Gln Lys -5 1 5

Ala Arg Val Ser Gln Glu Leu Lys Gly Ala Lys Lys Val His Leu Gly
10 15 20 25

Glu Asp Leu Lys Ser Ile Leu Ser Glu Ala Pro Gly Lys Cys Val Pro 30 35 40

Tyr Ala Val Ile Glu Gly Ala Val Arg Ser Val Lys Glu Thr Leu Asn 45 50 55

Ser Gin Phe Val Glu Asn Cys Xaa Gly Val Arg

60 65

- (2) INFORMATION FOR SEQ ID NO: 422:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 142 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide .
    - (B) LOCATION: -139..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.3 seq GILVPHSLRQAQA/SF
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 422:
- Met Ala Ala Leu Asp Leu Arg Ala Xaa Trp Ile Arg Trp Ser Cys Ser -135 -130
- Cys Leu Gly Xaa Leu Xaa Gly Ala Gly Glu Thr Asn Gly Val Glu -115
- Arg Pro Gly Gly Gly Leu Ala Leu Ala Arg Gln Gly Ser Leu Arg -105 -100
- Asp Gly Arg Gln Val Gly Arg Ala Pro Ala Val Cys Phe Pro His Gly
- Ala Pro Gly Leu Pro Pro Arg Gln Arg Xaa Xaa Gly Gly Xaa Pro Glu
- Val Gln Gly Glu Ser Trp Cys Pro Arg Pro Arg Gly Gly Ala
- Ser Arg Thr Gly Leu Arg Arg Lys Gly Pro Thr Lys Thr Pro Glu -40
- Pro Glu Ser Ser Glu Ala Pro Gln Asp Pro Leu Asn Trp Phe Gly Ile
- Leu Val Pro His Ser Leu Arg Gln Ala Gln Ala Ser Phe Arg -10
- (2) INFORMATION FOR SEQ ID NO: 423:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 29 amino acids
    - (B) TYPE: AMINO ACID

- (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -21..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.1

seg WWISLLPSLLSIC/KV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 423:

Met Ala Phe Leu Pro Ser Pro Ala Trp Trp Ile Ser Leu Leu Pro Ser
-20 -15 -10

Leu Leu Ser Ile Cys Lys Val Leu Met Pro Lys Leu Lys

- (2) INFORMATION FOR SEQ ID NO: 424:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 127 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -49..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.8

seq PAFHLPLPGPTLA/FL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 424:

Met Glu Pro Lys Val Ala Glu Leu Lys Gln Lys Ile Glu Asp Thr Leu
-45 -40 -35

Cys Pro Phe Gly Phe Glu Val Tyr Pro Phe Gln Val Ala Trp Tyr Asn
-30
-25
-20

Glu Leu Leu Pro Pro Ala Phe His Leu Pro Leu Pro Gly Pro Thr Leu
-15 -10 -5

Ala Phe Leu Val Leu Ser Thr Pro Ala Met Phe Asp Arg Ala Leu Lys
1 5 10 15

Pro Phe Leu Gln Ser Cys His Leu Arg Met Leu Thr Asp Pro Val Asp

20 25 30

Gln Cys Val Ala Tyr His Leu Gly Arg Val Arg Glu Ser Leu Pro Glu 35 40 45

Leu Gln Ile Glu Ile Ile Ala Xaa Xaa Arg Gly Ala Pro Gln Pro Thr
50 55 60

Pro Gln Asp Pro Gly Pro Asp Ser Ser His Val Ala Gly Ala Ala 65 70 75

# (2) INFORMATION FOR SEQ ID NO: 425:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 98 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -13..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.7

seq MLVLRSGLTKALA/SR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 425:

Met Leu Val Leu Arg Ser Gly Leu Thr Lys Ala Leu Ala Ser Arg Thr
-10 -5 1

Leu Ala Pro Gln Val Cys Ser Ser Phe Ala Thr Gly Pro Arg Gln Tyr
5 10 15

Asp Gly Thr Phe Tyr Glu Phe Arg Thr Tyr Tyr Leu Lys Pro Ser Asn 20 25 30 35

Met Asn Ala Phe Met Glu Asn Leu Lys Lys Asn Ile His Leu Arg Thr 40 45 50

Ser Tyr Ser Glu Leu Val Gly Phe Trp Ser Val Glu Phe Gly Gly Arg 55 60 65

Thr Asn Lys Val Phe His Ile Trp Lys Tyr Asp Asn Phe Ala His Arg
70 75 80

Ala Glu 85

(2) INFORMATION FOR SEQ ID NO: 426:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 27 amino acids

(B) TYPE: AMINO ACID (D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: -23..-1

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 10.5

seq LLFVLLLFSLLPA/CL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 426:

Met Ser Gly Gly His Leu Ala Asp Leu Thr Leu Leu Phe Val Leu Leu -20 -15 -10

Leu Phe Ser Leu Leu Pro Ala Cys Leu Pro Arg
-5 . 1

- (2) INFORMATION FOR SEQ ID NO: 427:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 54 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -51..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 10.1

seq LLGALTLLGLVTS/FY

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 427:

Met Lys Pro Ser Arg Thr Pro Ala Arg Leu Trp Met Leu Pro Gln Gln -50 -45 -40

Gln Ala Gly Ala Val Val Val Ala Ala Pro Thr Glu Arg His Pro Thr
-35 -25 -20

His His Met Ala Gly Trp Leu Leu Gly Ala Leu Thr Leu Leu Gly Leu
-15 -10 -5

Val Thr Ser Phe Tyr Lys

- (2) INFORMATION FOR SEQ ID NO: 428:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 131 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -64..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 9.5

seq LSLLAALAHLAAA/EK

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 428:
- Met Gly Glu Ser Ile Pro Leu Ala Ala Pro Val Pro Val Glu Gln Ala
  -60 -55 -50
- Val Leu Glu Thr Phe Phe Ser His Leu Gly Ile Phe Ser Tyr Asp Lys
  -45 -40 -35
- Ala Lys Asp Asn Val Glu Lys Glu Arg Glu Ala Asn Lys Ser Ala Gly
  -30 -25 -20
- Gly Ser Trp Leu Ser Leu Leu Ala Ala Leu Ala His Leu Ala Ala Ala -15 -10 -5
- Glu Lys Val Tyr His Ser Leu Thr Tyr Leu Gly Gln Lys Leu Gly Gly
  1 10 15
- Gln Ser Phe Phe Ser Arg Lys Asp Ser Ile Arg Thr Ile Tyr Thr Ser 20 25 30
- Leu His Asn Glu Leu Lys Lys Val Val Thr Gly Arg Gly Ala Xaa Xaa 35 40 45
- Trp Asp Cys Ser Ser Arg Gly Arg Thr Pro Phe Pro Pro Val Arg Ala 50 55 60

Ala Tyr Gly 65

- (2) INFORMATION FOR SEQ ID NO: 429:
  - (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 86 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig peptide
  - (B) LOCATION: -38..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 9.3

seq QLLYLSLLSGLHG/QE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 429:

Met Gln Met Ser Tyr Ala Ile Arg Cys Ala Phe Tyr Gln Leu Leu Leu -35 -30 -25

Ala Ala Leu Met Leu Val Ala Met Leu Gln Leu Leu Tyr Leu Ser Leu
-20 -15 -10

Leu Ser Gly Leu His Gly Gln Glu Glu Gln Asp Gln Tyr Phe Glu Phe
-5
1
5
10

Phe Pro Pro Ser Pro Arg Ser Val Asp Gln Val Lys Ala Gln Leu Arg
15 20 25

Thr Ala Leu Ala Ser Gly Gly Val Leu Asp Ala Ser Gly Asp Tyr Arg 30 35 40

Val Tyr Arg Gly His Gly
45

- (2) INFORMATION FOR SEQ ID NO: 430:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 26 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -24..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 9.1

seq LFAFHLLLSFILG/SR

(Mi) SEQUENCE DESCRIPTION: SEQ ID NO: 430:

Met Leu Arg Ala Glu Leu Lys Ile Ala Val Leu Phe Ala Phe His

Leu Leu Ser Phe Ile Leu Gly Ser Arg
-5 1

- (2) INFORMATION FOR SEQ ID NO: 431:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 76 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -55..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 9

seq LLILLLRTFLCSA/MI

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 431:

Met Asn His Gln Gln Thr Leu Ile Gly Arg Leu Leu Cys Asp Leu His
-55
-45
-40

Gly Leu Ser Leu Ser Pro Pro Val Ala Asn Asn Val Gln Ala Leu Phe
-35
-30
-25

Arg Met Leu Thr Pro Glu Ala Tyr Ser Cys Leu Leu Ile Leu Leu Leu -20 -15 -10

Arg Thr Phe Leu Cys Ser Ala Met Ile Ala Asn Thr Leu His Leu Lys
-5 1 5

Tyr His Leu Gln Leu Ile Asp Asn Ala Cys Pro Glu 10 15 20

- (2) INFORMATION FOR SEQ ID NO: 432:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 84 amino acids
    - (3) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: -40..-1

- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 8.6

seq LFCVLGIVLLVTG/IV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 432:

Met Ile Ile Thr Ala Val Val Ser Ile Ser Val Thr Ile Phe Cys Phe
-40 -35 -30 -25

Gln Thr Lys Val Asp Phe Thr Ser Cys Thr Gly Leu Phe Cys Val Leu
-20
-15
-10

Gly Ile Val Leu Leu Val Thr Gly Ile Val Thr Ser Ile Val Leu Tyr
-5
1
5

Phe Gln Tyr Val Tyr Trp Leu His Met Leu Tyr Ala Ala Leu Gly Ala 10 20

Ile Cys Phe Thr Leu Phe Leu Ala Tyr Asp Thr Gln Leu Val Leu Gly 25 30 35 40

Asn Arg Lys His

- (2) INFORMATION FOR SEQ ID NO: 433:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 89 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -65..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 8.4

seq LLWFIHLVFVVLX/LF

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 433:

Met Ala Ala Gly Gly Arg Met Glu Asp Gly Ser Leu Asp Ile Thr Gln -65 -55 -50

Ser Ile Glu Asp Asp Pro Leu Leu Asp Ala Gln Leu Leu Pro His His -45 -40 -35

Ser Leu Gln Ala His Phe Arg Pro Arg Phe His Pro Leu Pro Thr Val

. -30 -25 -20

Ile Ile Val Asn Leu Leu Trp Phe Ile His Leu Val Phe Val Val Leu
-15 -10 -5

Xaa Leu Phe Asn Arg Cys Ala Leu Phe Xaa Ser Tyr Pro Lys Trp Asp 1 5 10 15

Xaa Cys Pro Gly Asn Tyr Thr Asn Pro

- (2) INFORMATION FOR SEQ ID NO: 434:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 59 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -16..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7.5

seq GCMLLFVFGFVGG/AV

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 434:
- Met Ser Pro Gly Cys Met Leu Leu Phe Val Phe Gly Phe Val Gly Gly
  -15 -5
- Ala Val Val Ile Asn Ser Ala Ile Leu Val Ser Leu Ser Val Leu Leu 1 5 10 15
- Leu Val His Phe Ser Ile Ser Thr Gly Val Pro Ala Leu Thr Gln Asn 20 25 30

Leu Pro Arg Ile Leu Arg Lys Glu Arg Pro Gly 35

- (2) INFORMATION FOR SEQ ID NO: 435:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 67 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -15..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 7.5

seq LLLGIALLAYVAS/VW

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 435:

Met Lys Leu Leu Gly Ile Ala Leu Leu Ala Tyr Val Ala Ser Val -15 -10 -5

Trp Gly Asn Pne Val Asn Met Arg Ser Ile Gln Glu Asn Gly Glu Leu
5 10 15

Lys Ile Glu Ser Lys Ile Glu Glu Met Val Glu Pro Leu Arg Glu Lys 20 25 30

Ile Arg Asp Leu Xaa Lys Ser Phe Thr Gln Lys Tyr Pro Pro Val Lys 35 40 45

Phe Leu Ser 50

- (2) INFORMATION FOR SEQ ID NO: 436:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 151 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -23..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7.5

seq LVLLLTLPLHLMA/LL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 436:

Met Asp Ile Leu Val Pro Leu Leu Gln Leu Leu Val Leu Leu Leu Thr
-20 -15 -10

Leu Pro Leu His Leu Met Ala Leu Leu Gly Cys Trp Gln Pro Leu Cys
-5 5

Lys Ser Tyr Phe Pro Tyr Leu Met Ala Val Leu Thr Pro Lys Ser Asn 10 15 20 25 Arg Lys Met Glu Ser Lys Lys Arg Glu Leu Phe Ser Gln Ile Lys Gly  $30 \hspace{1cm} 35 \hspace{1cm} 40$ 

Leu Thr Gly Ala Ser Gly Lys Val Ala Leu Leu Glu Leu Gly Cys Gly
45 50 55

Thr Gly Ala Asn Phe Gln Phe Tyr Pro Pro Gly Cys Arg Val Thr Cys
60 70

Leu Asp Pro Asn Pro His Phe Glu Lys Phe Leu Thr Lys Ser Met Ala 75 80 85

Glu Asn Arg His Leu Gln Tyr Glu Arg Phe Val Val Ala Pro Gly Glu 90 95 100 105

Asp Met Arg Xaa Leu Ala Asp Gly Ser Met Asp Val Val Cys Thr

Leu Val Leu Cys Ser Val Gln 125

## (2) INFORMATION FOR SEQ ID NO: 437:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 103 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -35..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 7.4 seq SLLLSLELASGSG/QG
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 437:

Met Glu Ala Ala Ser Pro Ser Asn Ser Thr Gly Val Glu Arg Xaa Ala -35 -20 -25

Asp Leu Met Asp Ala Asp Ser Leu Leu Leu Ser Leu Glu Leu Ala Ser
-15 -10 -5

Gly Ser Gly Gln Gly Leu Ser Pro Asp Arg Arg Ala Ser Leu Leu Thr  $1 \hspace{1cm} 5 \hspace{1cm} 10$ 

Ser Leu Met Leu Val Lys Arg Asp Tyr Arg Tyr Asp Arg Val Leu Phe
15 20 . 25

Trp Gly Arg Ile Leu Gly Leu Val Ala Asp Tyr Tyr Ile Ala Gln Gly 30 40 45

Leu Ser Glu Asp Gln Leu Ala Pro Arg Lys Thr Leu Tyr Arg Ser Arg
50 55 60

Ser Arg Lys Arg Pro Ala Leu 65 \*

- (2) INFORMATION FOR SEQ ID NO: 438:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 70 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -43..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7.4

seq VLVKLLSSSASTS/RP

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 438:
- Met Ile Arg Gln Glu Arg Ser Thr Ser Tyr Gln Glu Ala Val Arg Pro
  -40 -35 -30
- Ala Leu Pro Ser Ser Lys Pro Cys Leu Leu Thr Ser Pro Ala Val Leu
  -25
  -20
  -15
- Val Lys Leu Leu Ser Ser Ser Ala Ser Thr Ser Arg Pro Pro Asp Leu
  -10 -5 1 5
- Gly His Leu Trp Gln Pro Ser Ser Ser Val Pro Leu His Arg Pro Pro 10 15 20

His Thr Ala Pro Pro Ala 25

- (2) INFORMATION FOR SEQ ID NO: 439:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 113 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain

- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -41..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 7.4

seq ILPLLFGCLGVFG/LF

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 439:

Met Lys Leu Ile Asp Tyr Gly Leu Ser Gly Tyr Gln Glu Glu Ser Ala
-40 -35 -30

Glu Val Lys Ala Met Asp Phe Ile Thr Ser Thr Ala Ile Leu Pro Leu -25 -15 -10

Leu Phe Gly Cys Leu Gly Val Phe Gly Leu Phe Arg Leu Leu Gln Trp -5 1 5

Val Arg Gly Lys Ala Tyr Leu Arg Asn Ala Val Val Val Ile Thr Gly
10 15 20

Ala Thr Ser Gly Leu Gly Lys Glu Cys Ala Lys Val Phe Tyr Ala Xaa 25 30 35

Gly Ala Lys Leu Val Leu Cys Glu Xaa Glu Trp Trp Gly Leu Glu Glu 40 55 55

Leu Ile Arg Glu Leu Thr Ala Ser His Ala Thr Lys Val Gln Thr His  $\phantom{-}60\phantom{+}65\phantom{+}70\phantom{+}$ 

Lys

- (2) INFORMATION FOR SEQ ID NO: 440:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 44 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -19..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7.4

seq PMLLRALAQAARA/GP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 440:

Met Arg Cys Leu Thr Thr Pro Met Leu Leu Arg Ala Leu Ala Gln Ala
-15
-10
-5

Ala Arg Ala Gly Pro Pro Gly Gly Arg Ser Leu His Ser Ser Ala Val

Ala Ala Thr Tyr Lys Tyr Val Asn Met Gln Asp Gln
15 20 25

- (2) INFORMATION FOR SEQ ID NO: 441:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 84 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -67..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 7

seq IWTLLSSVIRCLC/AI

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 441:
- Met Ser Arg Phe Leu Asn Val Leu Arg Ser Trp Leu Val Met Val Ser
  -65 -60 -55
- Ile Ile Ala Met Gly Asn Thr Leu Gln Ser Phe Arg Asp His Thr Phe  $\sim 50$   $\sim 45$   $\sim 40$
- Leu Tyr Glu Lys Leu Tyr Thr Gly Lys Pro Asn Leu Val Asn Gly Leu -35 -25 -20
- Gin Ala Arg Thr Phe Gly Ile Trp Thr Leu Leu Ser Ser Val Ile Arg
  -15 -10 -5
- Cys Leu Cys Ala Ile Asp Ile His Asn Lys Thr Leu Tyr His Ile Thr 1 5 10

Leu Trp Thr Phe 15

- (2) INFORMATION FOR SEQ ID NO: 442:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 29 amino acids
    - (E) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo Sapiens
- (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -14..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 6.8

seq IFLTLSLDSRVSA/IR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 442:

Met Ile Phe Leu Thr Leu Ser Leu Asp Ser Arg Val Ser Ala Ile Arg
-10 -5 1

Ser Pro Asn Phe Val Tyr Arg Ser Pro Thr Xaa His Gly 5 10 15

- (2) INFORMATION FOR SEQ ID NO: 443:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 101 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -29..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 6.8

seq LIFLCGAALLXVG/IW

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 443:
- Met Gln Cys Phe Ser Phe Ile Lys Thr Met Met Ile Leu Phe Asn Leu
  -25 -20 -15
- Leu Ile Phe Leu Cys Gly Ala Ala Leu Leu Xaa Val Gly Ile Trp Val
- Ser Ile Asp Gly Ala Ser Phe Leu Lys Ile Phe Gly Pro Leu Ser Ser 5 10 15
- Ser Ala Met Gln Phe Val Asn Val Gly Tyr Phe Leu Ile Ala Ala Gly 20 25 30 35
- Val Val Phe Ala Leu Gly Phe Leu Gly Cys Tyr Xaa Ala Lys Thr 40 45 50
- Glu Ser Xaa Cys Ala Leu Val Thr Phe Phe Xaa Ile Leu Leu Ile 55 60 65

Phe Ile Ala Asp Val

(2) INFORMATION FOR SEQ ID NO: 444:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 97 amino acids
- (B) TYPE: AMINO ACID
- (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -35..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 6.4

seq SACLLLCPTWTNP/QL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 444:

Met Ala Glu Ala Ala Leu Glu Ala Val Arg Xaa Ser Tyr Glu Asn Ser -35 -25 -20

Arg Pro Leu Gln Gly Ser Ser Ala Cys Leu Leu Cys Pro Thr Trp
-15
-10
-5

Thr Asn Pro Gln Leu Arg Ser Thr Ser Thr Gly Thr Gly Ser Ala Pro
1 5 10

Thr Gly Arg Ala Leu Ser Ala Thr Leu Cys Ser Thr Gly Arg Pro Ser 15 20 25

Xaa Xaa Trp Ser Leu Pro Tyr Phe Arg Ala Thr Val Gly Ser Thr Glu 30 40 45

Val Ser Val Ala Val Thr Pro Asp Gly Tyr Ala Asp Ala Val Arg Xaa 50 55 60

Asp

- (2) INFORMATION FOR SEQ ID NO: 445:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 34 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -19..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 6.4

seq SVFLLMVNGQVES/AQ

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 445:

Met Ala Thr Ala Ser Pro Ser Val Phe Leu Leu Met Val Asn Gly Gln
-15 -10 -5

Val Glu Ser Ala Gln Phe Pro Glu Tyr Asp Asp Leu Tyr Cys Lys Tyr  $1 \hspace{1cm} 5 \hspace{1cm} 10$ 

Cys Gln 15

- (2) INFORMATION FOR SEQ ID NO: 446:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 36 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -28..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 6

seq IGLMFLMLGCALP/IY

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 446:

Met Ala Gly Ile Lys Ala Leu Ile Ser Leu Ser Phe Gly Gly Ala Ile
-25 -20 -15

Gly Leu Met Phe Leu Met Leu Gly Cys Ala Leu Pro Ile Tyr Asn Lys
-10 -5

Tyr Trp Pro Trp

- (2) INFORMATION FOR SEQ ID NO: 447:
  - (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 39 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: -17..-1

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 5.9

seq ILLFGTLLMNAGA/VL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 447:

Met Ile Gly Asp Ile Leu Leu Phe Gly Thr Leu Leu Met Asn Ala Gly
-15 -10 -5

Ala Val Leu Asn Phe Lys Leu Lys Lys Lys Asp Thr Gln Gly Phe Gly  $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$ 

Glu Glu Ser Arg Glu Pro Trp

- (2) INFORMATION FOR SEQ ID NO: 448:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 28 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -16..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.9

seq MILTLSLFGSCIS/NF

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 448:

Met Lys Thr Met Ile Leu Thr Leu Ser Leu Phe Gly Ser Cys Ile Ser -15 -5

Asn Phe Glu Arg Tyr Met Thr Glu Arg Ser Ile Gln 1 5 10

- (2) INFORMATION FOR SEQ ID NO: 449:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 82 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -39..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.9

seq SVSVLSSLGIVLA/VV

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 449:
- Met Asp Trp Arg Val Pro Pro Ser Xaa Xaa Asp Pro Gly His Gln Asp
  -35
  -30
  -25
- Ile Pro Leu Pro Val Thr Xaa Xaa Phe Ile Ser Val Ser Val Leu Ser
  -20 -15 -10
- Ser Leu Gly Ile Val Leu Ala Val Val Cys Leu Ser Phe Asn Ile Tyr
  -5 1 5
- Asn Ser His Val Arg Tyr Ile Gln Asn Ser Gln Pro Asn Leu Asn Asn 10 20 25
- Leu Thr Ala Val Gly Cys Ser Xaa Ala Leu Ala Ala Val Phe Pro Trp 30 35 40

Gly Ser

- (2) INFORMATION FOR SEQ ID NO: 450:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 113 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -16..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.8

# seq AALPAWLSLQSRA/RT

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 450:

Met Ala Ala Ala Leu Pro Ala Trp Leu Ser Leu Gln Ser Arg Ala
-15
-10
-5

Arg Thr Leu Arg Ala Phe Ser Thr Ala Val Tyr Ser Ala Thr Pro Val
1 5 10 15

Pro Xaa Pro Ser Leu Pro Glu Arg Thr Pro Gly Asn Glu Arg Pro Pro 20 25 30

Arg Arg Lys Ala Leu Pro Pro Arg Thr Glu Lys Met Ala Val Asp Gln 35 40 45

Asp Trp Pro Xaa Val Tyr Pro Val Ala Ala Pro Phe Lys Pro Ser Ala 50 55 60

Val Pro Leu Pro Val Arg Met Gly Tyr Pro Val Lys Lys Gly Val Pro 65 70 75 80

Trp Xaa Arg Arg Glu Ser Xaa Thr Phe Lys Asp Ser Asn Phe Leu His
85 90 95

Leu

- (2) INFORMATION FOR SEQ ID NO: 451:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 29 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -25..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.8 seq LWISACAMLLCHG/SL
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 451:

Met Ala Met Val Ser Ala Met Ser Trp Val Leu Tyr Leu Trp Ile Ser -25 -15 -10

Ala Cys Ala Met Leu Leu Cys His Gly Ser Leu Gln Arg
-5

- (2) INFORMATION FOR SEQ ID NO: 452:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 78 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -69..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.7

seq LCRLLCLVRLFCC/SS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 452:

Met Gly Lys Glu Trp Gly Trp Gln Glu Met Glu Asn Gly Gly Ala Ala
-65 -60 -55

Pro Ala Tro Gly Ala Gly Pro Pro Val His Pro Ala Pro Pro Val -50 -45 -40

Glu Lys Thr Leu Ser Trp Gly Cys Gly Phe Gly Leu His Ser Gly Phe
-35 -30 -25

Gly Gly Ser Gly Gly Val Gly Leu Cys Arg Leu Leu Cys Leu Val -20 -15 -10

Arg Leu Phe Cys Cys Ser Ser Ile Leu Tyr Gln Arg Gln Gly

- (2) INFORMATION FOR SEQ ID NO: 453:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 65 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -21..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.6 seq LVLSLQFLLLSYD/LF
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 453:

Met Leu Gln Thr Ser Asn Tyr Ser Leu Val Leu Ser Leu Gln Phe Leu
-20 -15 -10

Leu Leu Ser Tyr Asp Leu Phe Val Asn Ser Phe Ser Glu Leu Leu Gln -5 1 5 10

Lys Thr Pro Val Ile Gln Leu Val Leu Phe Ile Ile Gln Asp Ile Ala 15 20 25

Val Leu Phe Asn Ile Ile Ile Ile Phe Leu Met Phe Phe Asn Thr Ser 30 40

Arg

- (2) INFORMATION FOR SEQ ID NO: 454:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 27 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -24..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.5

seq MGVCLLIPGLATA/CI

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 454:

Met Trp Phe Glu Ile Leu Pro Gly Leu Ser Val Met Gly Val Cys Leu -20 -15 -10

Leu Ile Pro Gly Leu Ala Thr Ala Cys Ile Arg

- (2) INFORMATION FOR SEQ ID NO: 455:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 46 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain

- (ix) FEATURE:
  - (A) NAME/KEY: sig peptide
  - (B) LOCATION: -22..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5.5

seq LADPLXLFPFSEG/LP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 455:

Met Arg Pro Ser Pro Leu Ser Gly Ile Leu Ala Asp Pro Leu Xaa Leu
-20 -15 -10

Phe Pro Phe Ser Glu Gly Leu Pro Arg Arg Arg Ala Ala Ser Arg Ser
-5 1 5 10

Arg Leu Gln Thr Pro Ser Ala Arg Cys Ser Pro Arg Pro Gly
15 20

- (2) INFORMATION FOR SEQ ID NO: 456:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 30 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (E) LOCATION: -27..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.4

seq SLMMAQXFIPAVA/KV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 456:

Met Arg Glu Ser Leu Ser Xaa Arg Ser Trp His Leu Pro Ala Ser Leu
-25 -20 -15

Met Met Ala Gln Xaa Phe Ile Pro Ala Val Ala Lys Val Gly
-10 -5 1

- (2) INFORMATION FOR SEQ ID NO: 457:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 60 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(vi) ORIGINAL SOURCE:

- (A) NAME/KEY: sig\_peptide
- (B) LOCATION: -58..-1
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 5.4

seq LSLHLLATRACYG/IL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 457:

Met Ser Gly Val Val Pro Thr Ala Pro Glu Gln Pro Ala Xaa Glu Met -55 -50

Glu Asn Gln Thr Lys Pro Pro Asp Pro Arg Pro Asp Ala Pro Pro Glu

Tyr Ser Ser His Xaa Phe Thr Arg Thr Pro Trp Lys Gln Leu Ser Leu

His Leu Leu Ala Thr Arg Ala Cys Tyr Gly Ile Leu

- (2) INFORMATION FOR SEQ ID NO: 458:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 83 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -77..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.3

seq TWVFTCLVFFCFG/LS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 458:

Met Trp Arg Tyr Gln Phe Gly Trp Gly Val Ile Thr Arg Gly Pro Arg -70

Glu Ile Pro Phe Pro Pro Ser Leu Leu Ala Ser Glu Ser Leu Leu Pro -55 ·

Pro Leu Pro Asp Leu Val Leu Thr Cys Thr Ser Leu Gly Phe Val Thr

Arg Val Trp. Met Ser Leu Asn Leu Asn Glu Leu Ser Leu Tyr Ser Arg

Thr Trp Val Phe Thr Cys Leu Val Phe Phe Cys Phe Gly Leu Ser Xaa -10 -5 1

Ser Leu Gly

- (2) INFORMATION FOR SEQ ID NO: 459:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 39 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -29..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.2

seq FFMLLGSLLPVKI/IE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 459:

Met Val Lys Leu Val Ala Lys Ile Leu Cys Met Val Gly Val Phe
-25 -20 -15

Phe Phe Met Leu Leu Gly Ser Leu Leu Pro Val Lys Ile Ile Glu Thr -10 -5 1

Asp Phe Glu Lys Ala Pro Gly

- (2) INFORMATION FOR SEQ ID NO: 460:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 22 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -17..-1

(C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 5.2

seq IMCLIGLKANASS/ET

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 460:

Met Pro Val Ser Ile Met Cys Leu Ile Gly Leu Lys Ala Asn Ala Ser

Ser Glu Thr His Ser Gly

- (2) INFORMATION FOR SEQ ID NO: 461:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 26 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -17..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.2

seq LLYLVLEKLVSRA/FQ

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 461:

Met Lys Val Ile Leu Leu Tyr Leu Val Leu Glu Lys Leu Val Ser Arg

Ala Phe Gln Asn Val Glu Ala Pro His Gly
1 5

- (2) INFORMATION FOR SEQ ID NO: 462:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 46 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig peptide

- (B) LOCATION: -19..-1
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 5.1

seq LLLGGRVCXPSLA/VG

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 462:

Met Ala Val Thr Leu Ser Leu Leu Gly Gly Arg Val Cys Xaa Pro

Ser Leu Ala Val Gly Ser Arg Pro Gly Gly Trp Arg Ala Gln Ala Leu  $1 \hspace{1.5cm} 5 \hspace{1.5cm} 10$ 

Leu Ala Gly Ser Arg Thr Pro Ile Pro Thr Gly Asn Arg Arg
15 20 25

- (2) INFORMATION FOR SEQ ID NO: 463:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 49 amino acids
    - (3) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -23..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 5.1

seg LLPELGVVTPAQG/PR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 463:

Met Leu Asn Gln Thr Ser Gly Arg Thr Ser Leu Leu Pro Glu Leu Gly

Val Val Thr Pro Ala Gln Gly Pro Arg Arg Val Trp Cys Gly His

Ser Lys Ala Lys Ala Arg Lys Ser Tyr Cys Ala Arg Ala Ile Asp Cys 10 20 25

Gln

- (2) IMFORMATION FOR SEQ ID NO: 464:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 135 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR

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- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -79..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 5

seq SFLGFSAPTPIQA/LT

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 464:
- Met Thr Ser Glu Asn Leu Val Gln Thr Ala Pro Lys Lys Lys Asn
- Lys Gly Lys Lys Gly Leu Glu Pro Ser Gln Ser Thr Ala Ala Lys Val
- Pro Lys Lys Ala Lys Thr Trp Ile Pro Glu Val His Asp Gln Lys Ala
- Asp Val Ser Ala Trp Lys Asp Leu Phe Val Pro Arg Pro Val Leu Arg
- Ala Leu Ser Phe Leu Gly Phe Ser Ala Pro Thr Pro Ile Gln Ala Leu
- Thr Leu Ala Pro Ala Ile Arg Asp Lys Leu Asp Ile Leu Gly Ala Ala
- Glu Thr Gly Ser Gly Lys Thr Leu Ala Phe Ala Ile Pro Met Ile His 20
- Ala Val Leu Gln Trp Gln Lys Arg Asn Ala Ala Pro Pro Pro Ser Asn
- Thr Glu Ala Pro Pro Gly Glu
- (2) INFORMATION FOR SEQ ID NO: 465:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 79 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig peptide

- · (B) LOCATION: -23..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.9 seq WHXLIPLTWACMA/RQ
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 465:

Leu Thr Trp Ala Cys Met Ala Arg Gln Thr Pro His Leu Gly Glu Gln -5 5

Arg Arg Thr Thr Ala Ser Leu Xaa Arg Lys Leu Thr Thr Ala Ser Asn 10 20 25

Gly Gly Val Ile Glu Glu Leu Ser Cys Val Arg Ser Asn Asn Tyr Val 30 35 40

Gln Glu Pro Glu Cys Arg Arg Asn Leu Val Gln Cys Leu Leu Trp 45 50 55

- (2) INFORMATION FOR SEQ ID NO: 466:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 68 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -57..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.9 seq GWFLSGCPHGSSA/TW
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 466:

Met Ser Leu Thr Ser Ser Pro Lys Lys Arg Arg Ser Ile Cys Phe Asp
-55 -50 -45

Arg Phe Leu Met Pro Gln Ser Gln Ser Gly Pro Ser Ser Leu Gly Glu
-40 -35 -30

Ser Tyr Arg Thr Gly Val Gly Phe Leu Ile Pro Glu Gly Trp Phe Leu -25 -15 -10

Ser Gly Cys Pro His Gly Ser Ser Ala Thr Trp Thr Lys Cys Gln Thr
-5 1 5

Ser Ala Ser Leu

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10

```
(2) INFORMATION FOR SEQ ID NO: 467:
```

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 46 amino acids
- (B) TYPE: AMINO ACID
- (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -39..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix

415

(D) OTHER INFORMATION: score 4.8

seq SLXFCLSPPPSPS/LR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 467:

Met Gly Glu Leu Gly Asn Arg Ser Arg Cys Ile Leu Phe Leu Ser Glu
-35 -30 -25

Asn Pro Cys Leu Ser Glu Ser Ile Phe Gln Ser Leu Xaa Phe Cys Leu
-20 -15 -10

Ser Pro Pro Pro Ser Pro Ser Leu Arg Pro Ser Pro Ser Arg
-5 1 5

- (2) INFORMATION FOR SEQ ID NO: 468:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 111 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig peptide
    - (B) LOCATION: -93..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.7

seq VLLLRQXFAQAEK/WY

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 468:

Met Ala Glu Leu Gly Leu Asn Glu His His Gln Asn Glu Val Ile Asn
-90 -85 -80

Tyr Met Arg Phe Ala Arg Ser Lys Arg Gly Leu Arg Leu Lys Thr Val -75 -70 -65

Asp Ser Cys Phe Gln Asp Leu Lys Glu Ser Arg Leu Val Glu Asp Thr
-60 -55 -50

Phe Thr Ile Asp Glu Val Ser Glu Val Leu Asn Gly Leu Gln Ala Val
-45 -35 -30

Val His Ser Glu Val Glu Ser Glu Leu Ile Asn Thr Ala Tyr Thr Asn
-25
-20
-15

Val Leu Leu Arg Gln Xaa Phe Ala Gln Ala Glu Lys Trp Tyr Leu
-10 -5

Lys Leu Gln Thr Asp Ile Ser Glu Leu Glu Asn Arg Glu Leu Leu  $5 \hspace{1cm} 10 \hspace{1cm} 15$ 

### (2) INFORMATION FOR SEQ ID NO: 469:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 70 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -49..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.6

seq SWAVGLLYAVAQG/SK

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 469:

Met Val Thr Leu Pro Ser Gly Thr Trp Ala Phe Ser Cys Pro Tyr Leu
-45 -40 -35

Ala Leu Val Asp Gly Gly Met Leu Gly Ser Ala Arg Glu Asp Ala His

Ala Ser Val Val Ser Trp Ala Val Gly Leu Leu Tyr Ala Val Ala Gln
-15 -10 -5

Gly Ser Lys Arg Arg Lys Val Gln Asp Val Lys Pro Leu Xaa Trp Ser 1 5 10

Arg Thr Gly Thr Leu Gly

## (2) INFORMATION FOR SEQ ID NO: 470:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 80 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -68..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.6

seq LPFSLVSMLVTQG/LV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 470:

Met Ala Ser Ala Ser Ala Arg Gly Asn Gln Asp Lys Asp Ala His Phe

Pro Pro Pro Ser Lys Gln Ser Leu Leu Phe Cys Pro Lys Xaa Xaa Leu

His Ile His Arg Ala Glu Ile Ser Lys Ile Met Arg Glu Cys Gln Glu

Glu Ser Phe Trp Lys Arg Ala Leu Pro Phe Ser Leu Val Ser Met Leu

Val Thr Gln Gly Leu Val Tyr Gln Gly Tyr Leu Ala Ala Asn Ser Arg

- (2) INFORMATION FOR SEQ ID NO: 471:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 71 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (3) LOCATION: -69..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.5

seq FILSLCVLCIVLT/TG

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 471:

Met Leu Leu Met Lys Ser Ile Leu Leu Lys Val Val Cys Val Leu Cys
-65 -60 -55

Ile Tyr Leu Lys Phe Lys Leu Met Ala Leu Ile Tyr Val Pro Asp Lys
-50 -45 -40

Asn Asn Thr Asn Asn Ile Leu Arg Tyr Asn His Asn Glu Ile Ser
-35 -30 -25

Ile Gly Ile Ser Val Gln Cys His Phe Ile Leu Ser Leu Cys Val Leu
-20 -15 -10

Cys Ile Val Leu Thr Thr Gly

- (2) INFORMATION FOR SEQ ID NO: 472:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 50 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -16..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.5

seq RLLLRRFLASVIS/RK

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 472:
- Met Ala Gln Arg Leu Leu Arg Arg Phe Leu Ala Ser Val Ile Ser

Arg Lys Pro Ser Gln Gly Gln Trp Pro Pro Leu Thr Ser Arg Ala Leu 1 5 10

Gln Thr Pro Xaa Cys Ser Xaa Gly Gly Leu Thr Val Thr Pro Asn Pro 20 25 30

Ser Arg

- (2) INFORMATION FOR SEQ ID NO: 473:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 103 amino acids

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- (B) TYPE: AMINO ACID
  (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -77..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.4

seq FEARIALLPLLQA/ET

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 473:

Met Ala Ala Ser Lys Val Lys Gln Asp Met Pro Pro Xaa Gly Gly Tyr
-75 -70 -65

Gly Pro Ile Asp Tyr Lys Arg Asn Leu Pro Arg Arg Gly Leu Ser Gly
-60 -55 -50

Tyr Ser Met Leu Ala Ile Gly Ile Gly Thr Leu Ile Tyr Gly His Trp -45 -35 -30

Ser Ile Met Lys Trp Asn Arg Glu Arg Arg Arg Leu Gln Ile Glu Asp
-25 -20 -15

Phe Glu Ala Arg Ile Ala Leu Leu Pro Leu Leu Gln Ala Glu Thr Asp
-10 -5 1

Arg Arg Thr Leu Gln Met Leu Arg Glu Asn Leu Glu Glu Glu Ala Ile 5 10

Ile Met Lys Asp Val Pro Gly 20 25

- (2) INFORMATION FOR SEQ ID NO: 474:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 77 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -54..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.3

seq LLSLAILSHISTP/GC

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 474:

Met Arg His Leu Val Thr Glu Glu Leu Phe Pro Cys Ser Asn Leu Glu
-50 -45 -40

Asp Val Val Glu Asp Asn Ser His Ser Tyr Phe Thr Leu Arg Ile Thr -35 -30 -25

Met Ala Cys Lys Gly Val Pro Ser Thr Leu Leu Ser Leu Ala Ile Leu
-20 -15 -10

Ser His Ile Ser Thr Pro Gly Cys Glu Trp His Val Ile Tyr Val Ser
-5 1 5 10

Ser Xaa Gly Leu Tyr Leu Val Val Glu Met Thr Asp Arg

- (2) INFORMATION FOR SEQ ID NO: 475:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 80 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -76..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.3

seq FRLLXVFAYGTYA/DY

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 475:

Met Ser Ala Glu Val Lys Val Thr Gly Gln Asn Gln Glu Gln Phe Leu -75 -65

Leu Leu Ala Lys Ser Ala Lys Gly Ala Ala Leu Ala Thr Leu Ile His -60 -55 -50 -45

Gln Val Leu Glu Ala Pro Gly Val Tyr Val Phe Gly Glu Leu Leu Asp
-40 -35 -30

Met Pro Asn Val Arg Glu Leu Ala Glu Ser Xaa Phe Ala Ser Thr Phe
-25
-20
-15

Arg Leu Leu Xaa Val Phe Ala Tyr Gly Thr Tyr Ala Asp Tyr Xaa Ala

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- (2) INFORMATION FOR SEQ ID NO: 476:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 45 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -34..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.3

seq QLFAFLNLLPVEA/DI

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 476:

Met Leu Leu Ser Ile Gly Met Leu Met Leu Ser Ala Thr Gln Val Xaa

Thr Ile Leu Xaa Val Gln Leu Phe Ala Phe Leu Asn Leu Leu Pro Val

Glu Ala Asp Ile Xaa Ala Tyr Asn Phe Glu Asn Ala Ser 5

- (2) INFORMATION FOR SEQ ID NO: 477:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 28 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -16..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.2

seg EVVSLSYCGVSWG/RI

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 477:

Met Gly Trp Glu Val Val Ser Leu Ser Tyr Cys Gly Val Ser Trp Gly -15 -10

Arg Ile Ser Pro Asn Leu Asn Lys Pro Val Asn Arg

1 . 5 10

(2) INFORMATION FOR SEQ ID NO: 478:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 72 amino acids
- (B) TYPE: AMINO ACID
- (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -32..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.2

seq CWELFCLEHGIQA/DG

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 478:

Met Arg Glu Cys Ile Ser Val His Val Gly Gln Ala Gly Val Gln Ile
-30 -25 -20

Gly Asn Ala Cys Trp Glu Leu Phe Cys Leu Glu His Gly Ile Gln Ala
-15 -10 -5

Asp Gly Thr Phe Asp Ala Gln Ala Ser Lys Ile Asn Asp Asp Ser 1 5 10 15

Phe Thr Thr Phe Phe Ser Glu Thr Gly Thr Ser Leu Leu Met Glu Arg 20 25 30

Leu Xaa Leu Asp Tyr Gly Lys Lys 35 40

- (2) INFORMATION FOR SEQ ID NO: 479:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 46 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -25..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4.2

seq LDLLRGLPRVSLA/NL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 479:

Met Ala Gly Pro Leu Gln Gly Gly Gly Ala Arg Ala Leu Asp Leu Leu -25 -10 -10

Arg Gly Leu Pro Arg Val Ser Leu Ala Asn Leu Lys Pro Asn Pro Gly
-5 1 5

Ser Lys Lys Pro Glu Arg Arg Pro Arg Gly Arg Arg Trp
10 15 20

- (2) INFORMATION FOR SEQ ID NO: 480:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 68 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -25..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.2 seq MFAASXLAMCAGA/EV
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 480:

Met Pro Ala Gly Val Pro Met Ser Thr Tyr Leu Lys Met Phe Ala Ala -25 -15 -10

Ser Xaa Leu Ala Met Cys Ala Gly Ala Glu Val Val His Arg Tyr Tyr -5 1 5

Arg Pro Asp Leu Thr Ile Pro Glu Ile Pro Pro Lys Arg Gly Glu Leu 10 15 20

Lys Thr Glu Leu Leu Gly Leu Lys Glu Arg Lys His Lys Pro Gln Val 25 30 35

Ser Gln Gln Glu

40

- (2) INFORMATION FOR SEQ ID NO: 481:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 82 amino acids

- (B) TYPE: AMINO ACID
  (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -24..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix

424

(D) OTHER INFORMATION: score 4.2

seq SLPALALSLRASP/RX

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 481:

Met Ala Val Gln Cys Val Arg Leu Ala Arg Arg Ser Leu Pro Ala Leu
-20 -15 -10

Ala Leu Ser Leu Arg Ala Ser Pro Arg Xaa Leu Cys Thr Ala Thr Lys
-5 5

Gln Lys Asn Ser Gly Gln Asn Leu Glu Glu Asp Met Gly Gln Ser Glu 10 20

Gln Lys Ala Asp Pro Pro Ala Thr Glu Lys Thr Leu Leu Glu Glu Lys 25 30 35 40

Val Lys Leu Glu Glu Glu Lys Glu Thr Val Glu Lys Tyr Lys Arg 45 50 55

Ala Arg

- (2) INFORMATION FOR SEQ ID NO: 482:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 42 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -37..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.2

seq RLMHHYLSTPTSA/RP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 482:

Met Phe Ser Ile Ile Ser Arg Ser Arg Ala Cys Ser Met Tyr Phe Lys
-35
-30
-25

Glu Asn Ala Lys Pro Ser Gln Leu Arg Leu Met His His Tyr Leu Ser
-20 -15 -10

Thr Pro Thr Ser Ala Arg Pro His His Leu
-5 1 5

- (2) INFORMATION FOR SEQ ID NO: 483:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 35 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -16..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 4.1

seq LLPATSLAGPVLS/TL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 483:

Met Lys Arg Leu Leu Pro Ala Thr Ser Leu Ala Gly Pro Val Leu Ser -15 -10 -5

Thr Leu Ile Ala Pro Thr Pro Met Leu Phe Cys Glu Asp Lys Ser Trp 1 5 10 15

Asp Pro Gly

- (2) INFORMATION FOR SEQ ID NO: 484:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 98 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -70..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix

(D) OTHER INFORMATION: score 4

seq IAVLYLHLYDVFG/DP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 484:

Met Leu Ile Ile Thr Asn Pro Trp Pro Lys Tyr Phe Asp Ala Ala Gly
-70 -65 -60 -55

Arg Leu Thr Pro Glu Phe Ser Gln Arg Leu Thr Asn Lys Ile Arg Glu -50 . -45 . -40

Leu Leu Gln Gln Met Glu Arg Gly Leu Lys Ser Ala Asp Xaa Xaa Asp
-35
-30
-25

Gly Thr Gly Tyr Thr Gly Trp Ala Gly Ile Ala Val Leu Tyr Leu His
-20 -15 -10

Leu Tyr Asp Val Phe Gly Asp Pro Ala Tyr Leu Gln Leu Ala His Gly
-5 1 5

Tyr Val Lys Gln Ser Leu Asn Cys Leu Thr Lys Arg Ser Ile Thr Phe 15 20 25

Gln Gly

### (2) INFORMATION FOR SEQ ID NO: 485:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 47 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide.
  - (B) LOCATION: -21..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.9 seq AWLAQGSSSAGWG/LE
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 485:

Met Cys Ala Thr Glu Thr Val Arg Ala Trp Leu Ala Gln Gly Ser Ser -20 -15 -10

Ser Ala Gly Trp Gly Leu Glu Arg Lys Gln Gly Val Ser Ala His Arg -5 10

Met Pro Ala Leu Arg Trp Leu Gln Lys Ser Val Pro Gly Xaa Met 15 20 25

(2) INFORMATION FOR SEQ ID NO: 486:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 48 amino acids
- (B) TYPE: AMINO ACID
- (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -46..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.9

seq AAAFCLKXXGANT/HP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 486:

Met Leu Leu Ala Thr His Pro Glu Thr Val Gly Gln Val Thr Leu
-45 -40 -35

Arg Val Xaa Pro Val Ser Leu Glu Val Ser Ile Gln Met Cys Ala Ala -30 -25 -20 -15

Ala Ala Ala Ala Phe Cys Leu Lys Xaa Xaa Gly Ala Asn Thr His Pro
-10 -5 1

- (2) INFORMATION FOR SEQ ID NO: 487:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 86 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -64..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.9

seq GLGGAQLQGGAXG/RG

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 487:

Met Ala Ala Ser Ser Ala Thr Pro Ala Pro Xaa Xaa Ser Gln Arg Cys
-60 -55 -50

Gly Ala Asp Ala Gly Ser Ala Ala Arg Ile Val Phe Arg Trp Gly Arg
-45 -40 -35

Gly Arg Arg Gly Ala Arg Ser Pro Glu Gly Ser Gly His His Gly Arg
-30 -25 -20

Ala Asn Ser Gly Leu Gly Gly Ala Gln Leu Gln Gly Gly Ala Xaa Gly
-15 -5

Arg Gly Ser Met Ala Pro Leu Arg Ala Ser Ala Gly Gln Thr Arg Asp

1 10 15

Gly Pro Thr Gln Pro Gly

- (2) INFORMATION FOR SEQ ID NO: 488:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 50 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (3) LOCATION: -17..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.9

seq PLAGLAAAALGRA/PP

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 488:

Met Leu Arg Arg Pro Leu Ala Gly Leu Ala Ala Ala Ala Leu Gly Arg
-15 -10 -5

Ala Pro Pro Asp Gly Leu Leu Cys Ser Leu Pro Gly Val Ala Val Glu
1 5 10 15

Asp Pro Val Gln Asp Ser Ala Gly Phe Ser Phe Ser Leu Met Asp Arg
20 25 30

Pro Lys

- (2) INFORMATION FOR SEQ ID NO: 489:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 53 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

(B) LOCATION: -17..-1

- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 3.8

seq GFVAALVAGGVAG/VS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 489:

Met Asp Arg Pro Gly Phe Val Ala Ala Leu Val Ala Gly Gly Val Ala
-15
-10
-5

Gly Val Ser Val Asp Leu Ile Leu Phe Pro Leu Asp Thr Ile Lys Thr 1 5 10 15

Arg Leu Gln Ser Pro Gln Gly Phe Ser Lys Ala Gly Gly Phe His Gly 20 25 30

Ile Tyr Ala Ser Trp

### (2) INFORMATION FOR SEQ ID NO: 490:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 50 amino acids
  - (B) TYPE: AMINO ACID
  - (D) TOPOLOGY: LINEAR
- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -21..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.8

seq SMDLLTLLFQRRS/HQ

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 490:

Met Ile Val Trp Phe Glu Gly Ile Ser Met Asp Leu Leu Thr Leu Leu
-20 -15 -10

Phe Gln Arg Arg Ser His Gln Val Thr Gln Leu Leu Val Ser Ser Thr -5 1 5 . 10

Gly Asn Trp Leu Arg Gln Tyr Leu Cys Ala Ser Leu Thr Ile Ala Gly
15 20 25

WO 99/06552

Arg Arg .

- (2) INFORMATION FOR SEQ ID NO: 491:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 50 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -20..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix

430

(D) OTHER INFORMATION: score 3.8

seq ALDALMFPARRRA/AV

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 491:

Met Arg Thr Phe Val His Phe Ala Leu Asp Ala Leu Met Phe Pro Ala -20 -15 -10 -5

Arg Arg Arg Ala Ala Val Thr Arg Leu Ser Glu Arg Leu Ser Leu Cys

1 5 10

Phe Cys Leu His Ser Arg Leu Gln Asp Pro Ala Ala Arg Pro Arg Pro
15 20 25

Ser Trp

- (2) INFORMATION FOR SEQ ID NO: 492:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 99 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -61..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.8

seq LVMTFLFRNGSLQ/EK

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 492:

Met Ala Ala Pro Pro Gln Leu Arg Ala Leu Leu Val Val Val Asn Ala
-60 -55 -50

Leu Leu Arg Lys Arg Arg Tyr His Ala Ala Leu Ala Val Leu Lys Gly
-45 -35 -30

Phe Arg Asn Gly Ala Val Tyr Gly Ala Lys Ile Arg Ala Pro His Ala
-25
-20
-15

Leu Val Met Thr Phe Leu Phe Arg Asn Gly Ser Leu Gln Glu Lys Leu
-10 -5 1

Trp Ala Ile Leu Gln Ala Thr Tyr Ile His Ser Trp Asn Leu Ala Arg 5 10 15

Phe Val Phe Thr Tyr Lys Gly Leu Arg Ala Leu Gln Ser Tyr Ile Gln 20 25 30 35

Gly Pro Gly

- (2) INFORMATION FOR SEQ ID NO: 493:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 79 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -18..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.8

seq GXALGLLPSLAKA/ED

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 493:

Met Pro Val Asp Leu Gly Xaa Ala Leu Gly Leu Leu Pro Ser Leu Ala
-15 -10 -5

Lys Ala Glu Asp Ser Gln Phe Ser Glu Ser Asp Ala Ala Leu Gln Glu  $1 \hspace{1.5cm} 5 \hspace{1.5cm} 10$ 

Glu Leu Ser Ser Pro Glu Thr Ala Arg Gln Leu Phe Arg Gln Phe Arg 15 20 25 30

Tyr Gln Val Met Ser Gly Pro His Glu Thr Leu Lys Xaa Leu Arg Lys 35 40 45

Leu Cys Phe Gln Trp Leu Gln Pro Glu Val His Thr Lys Glu Gly

50 55 60

- (2) INFORMATION FOR SEQ ID NO: 494:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 139 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -72..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.8

seq LMGLALAVYKCQS/MG

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 494:
- Met Asn Leu Phe Ile Met Tyr Met Ala Gly Asn Thr Ile Ser Ile Phe
  -70 -65 -60
- Pro Thr Met Met Val Cys Met Met Ala Trp Arg Pro Ile Gln Ala Leu
  -55 -45
- Met Ala Ile Ser Ala Thr Phe Lys Met Leu Glu Ser Ser Ser Gln Lys -40 -35 -30 -25
- Phe Leu Gln Gly Leu Val Tyr Leu Ile Gly Asn Leu Met Gly Leu Ala
  -20 -15 -10
- Leu Ala Val Tyr Lys Cys Gln Ser Met Gly Leu Leu Pro Thr His Ala
  -5 5
- Ser Asp Trp Leu Ala Phe Ile Glu Pro Pro Glu Arg Met Glu Ser Val 10 20
- Val Glu Asp Cys Phe Cys Glu His Glu Lys Ala Ala Pro Gly Pro Tyr 25 30 35 40
- Val Phe Gly Ser Tyr Leu His Pro Ser Leu Ser Pro Val Ala Pro Gln
  45 50 55
- His Thr Leu Lys Leu Ile Thr Tyr Val Lys Lys 60 65
- (2) INFORMATION FOR SEQ ID NO: 495:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 102 amino acids
    - (B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

(ii) MOLECULE TYPE: PROTEIN

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo Sapiens

(F) TISSUE TYPE: Brain

(ix) FEATURE:

(A) NAME/KEY: sig\_peptide

- (B) LOCATION: -51..-1
- (C) IDENTIFICATION METHOD: Von Heijne matrix
- (D) OTHER INFORMATION: score 3.8

seq NVLFVAGLAFVIG/LE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 495:

Met Ile Ser Leu Thr Asp Thr Gln Lys Ile Gly Met Gly Leu Thr Gly
-50 -45 -40

Phe Gly Val Phe Phe Leu Phe Phe Gly Met Ile Leu Phe Phe Asp Lys
-35
-25
-20

Ala Leu Leu Ala Ile Gly Asn Val Leu Phe Val Ala Gly Leu Ala Phe -15 -10 -5

Val Ile Gly Leu Glu Arg Thr Phe Arg Phe Phe Gln Lys His Lys

1 5 10

Met Lys Ala Thr Gly Phe Phe Leu Gly Gly Val Phe Val Val Leu Ile 15 20 25

Gly Trp Pro Leu Ile Gly Met Ile Phe Glu Ile Tyr Gly Phe Phe Leu 30 35 40 45

Leu Phe Arg Gly Leu Gly

- (2) INFORMATION FOR SEQ ID NO: 496:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 53 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -33..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.6

seq LAVFQMLKSMCAG/QR

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 496:

Met Ala Ala Ser Gly Ala Pro Arg Ile Leu Val Asp Leu Leu Lys Leu
-30
-25
-20

Xaa Val Ala Pro Leu Ala Val Phe Gln Met Leu Lys Ser Met Cys Ala
-15 -10 -5

Gly Gln Arg Leu Ala Ser Glu Pro Gln Asp Pro Ala Ala Val Ser Leu 1 5 10 15

Pro Thr Ser Ser Gly

- (2) INFORMATION FOR SEQ ID NO: 497:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 92 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -26..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.6

seq ARSLLQFLRLVGQ/LK

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 497:

Met Ala Ser Val Ser Ser Ala Thr Phe Ser Gly His Gly Ala Arg Ser
-25 -20 -15

Leu Leu Gln Phe Leu Arg Leu Val Gly Gln Leu Lys Arg Val Pro Arg
-10 -5 1 5

Thr Gly Trp Val Tyr Arg Asn Val Gln Arg Pro Glu Ser Val Ser Asp
10 15 20

His Met Tyr Arg Met Ala Val Met Ala Met Val Ile Lys Asp Asp Arg 25 30 35

Leu Asn Lys Asp Arg Cys Val Arg Leu Ala Leu Val His Asp Met Ala 40 45 50

Glu Cys Ile Val Gly Asp Ile Ala Pro Ala Asp Gly
55 60 65

(2) INFORMATION FOR SEQ ID NO: 498:

WO 99/06552 435

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 24 amino acids

(B) TYPE: AMINO ACID

(D) TOPOLOGY: LINEAR

- (ii) MOLECULE TYPE: PROTEIN
- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -16..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 3.6

seq LAVLLVLFTLNIL/KS

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 498:

Met Trp Tyr Leu Ala Val Leu Leu Val Leu Phe Thr Leu Asn Ile Leu
-15 -5

Lys Ser Leu Tyr Trp Gln Pro Gly

- (2) INFORMATION FOR SEQ ID NO: 499:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 58 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -42..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.5

seq INSLLEXSSLSRC/LE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 499:

Met Phe Thr Phe Gly Arg Leu Phe Gln Ile Ile Thr Val Val Thr Cys
-40 -35 -30

Leu Gln Phe Ile Gln Asp Cys Gys Ile His Ser Arg Gln Ile Asn Ser -25 -20 -15

Leu Leu Glu Xaa Ser Ser Leu Ser Arg Cys Leu Glu Val Pro Met Tyr
-10 -5 1 5

WO 99/06552

Val Lys Cys Ile Gly Ser Lys Ile Pro Leu 10 15

- (2) INFORMATION FOR SEQ ID NO: 500:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 62 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -51..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 3.5

seq VGTLCQLDWWIWG/GI

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 500:

Met Ile Gln Asp Arg Asp Arg Cys Ala Gln Ala Ala Ala Val Ala Ala -50 -45 -40

Val Gly Asn Leu Glu Pro Arg Gly Thr Pro Gly Pro Glu Asp Glu Ala
-35 -25 -20

Phe Cys Leu Pro Gly Cys Val Gly Thr Leu Cys Gln Leu Asp Trp Trp
-15 -10 -5

- (2) INFORMATION FOR SEQ ID NO: 501:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 48 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -31..-1
    - (C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 11.3

#### seq LLLCLLWIGYSQG/TT

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 501:

Met Lys Ile Ile Phe Pro Ile Leu Ser Asn Pro Val Phe Arg Arg Thr
-30 -25 -20

Val Lys Leu Leu Cys Leu Leu Trp Ile Gly Tyr Ser Gln Gly Thr
-15 -5 1

Thr His Val Leu Arg Phe Gly Gly Ile Phe Glu Tyr Val Glu Ser Gly 5 10 15

- (2) INFORMATION FOR SEQ ID NO: 502:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 57 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo Sapiens
    - (F) TISSUE TYPE: Brain
  - (ix) FEATURE:
    - (A) NAME/KEY: sig\_peptide
    - (B) LOCATION: -26..-1
    - ·(C) IDENTIFICATION METHOD: Von Heijne matrix
    - (D) OTHER INFORMATION: score 6.6

seq LFWLASGWTPAFA/YS

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 502:
- Met Val Ser Arg Met Val Ser Thr Met Leu Ser Gly Leu Leu Phe Trp
  -25 -15

Leu Ala Ser Gly Trp Thr Pro Ala Phe Ala Tyr Ser Pro Arg Thr Pro -10 -5 1 5

Asp Arg Val Ser Glu Ala Asp Ile Gln Arg Leu Leu His Gly Val Met
10 15 20

Glu Gln Leu Gly Ile Ala Arg Pro Arg

- (2) INFORMATION FOR SEQ ID NO: 503:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 46 amino acids
    - (B) TYPE: AMINO ACID
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN

- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo Sapiens
  - (F) TISSUE TYPE: Brain
- (ix) FEATURE:
  - (A) NAME/KEY: sig\_peptide
  - (B) LOCATION: -24..-1
  - (C) IDENTIFICATION METHOD: Von Heijne matrix
  - (D) OTHER INFORMATION: score 4.8

seq ATMVSGSSGLAXA/RL

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 503:

Met Thr Ala Thr Leu Ala Ala Ala Ala Asp Ile Ala Thr Met Val Ser -20 -15 -10

Gly Ser Ser Gly Leu Ala Xaa Ala Arg Leu Leu Ser Arg Xaa Ser Ser -5 1 5

Cys Arg Arg Met Glu Phe Gly Ile Val Pro Thr Gln Pro Arg 10 . 15 20

Internal Application No PCT / IB 98/01236

A. CLASSI IPC 6	FICATION OF SUBJECT MATTER C12N15/12 C07K14/47 C12N1	5/10 C12N15/11				
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Electronic d	ata base consulted during the international search (name of data	base and, where practical, search terms used				
C. DOCUM	ENTS CONSIDERED TO BE RELEVANT					
Category *	Citation of document, with indication, where appropriate, of the	relevant passages	Relevant to claim No.			
A	ADAMS M D ET AL: "3,400 NEW EXSEQUENCE TAGS IDENTIFY DIVERSITED TRANSCRIPTS IN HUMAN BRAIN" NATURE GENETICS, vol. 4, no. 3, July 1993, page: XP000645060 see the whole document	TY OF				
A	ADAMS M D ET AL: "RAPID CDNA!  (EXPRESSED SEQUENCE TAGS) FROM DIRECTIONALLY CLONED HUMAN INFO CDNA LIBRARY"  NATURE GENETICS,  vol. 4, no. 4, August 1993, pag STANDARD, XP002064427  see the whole document	ANT BRAIN				
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X Furt	her documents are listed in the continuation of box C.	X Patent family members are listed	in annex.			
1	stegories of cited documents : ent defining the general state of the lart which is not	"T" later document published after the inte or priority date and not in conflict with	rnational filing date the application but			
*E" earlier of filling of *L" docume	dereo to be of particular relevance document but published on or after the international	cited to understand the principle or th invention  "X" document of particular relevance; the cannot be considered novel or cannot involve an inventive step when the document.	claimed invention t be considered to cument is taken alone			
"O" docum other	no cither special reason (as specified) ent referring to an oral disclosure, use, exhibition or means ent published prior to the international filing date but	"Y" document of particular relevance; the c cannot be considered to involve an in document is combined with one or m ments, such combination being obvio in the art.	ventive step when the pre-other such docu- us to a person skilled			
later ti	nan the priority date claimed actual completion of the international search	"&" document member of the same patent  Date of mailing of the international ser				
	O November 1998	1 7. 02. 99	u en report			
Name and r	mailing address of the ISA  European Patent Office, P.B. 5818 Patentiaan 2  NL - 2280 HV Rijswijk  Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,  Fax: (+31-70) 340-3016	Authorized officer Smalt, R				

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Interretonal Application No
PC:/IB 98/01236

	•	PC1/1B 98/01236
C.(Continua	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	
Category 3	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Α	EP 0 279 582 A (BAYLOR COLLEGE MEDICINE) 24 August 1988 see the whole document	12,13
A	LIN Y ET AL: "INHIBITION OF NUCLEAR TRANSLOCATION OF TRANSCRIPTION FACTOR NF-KB BY A SYNTHETIC PEPTIDE CONTAINING A CELL MEMBRANE-PERMEABLE MOTIF AND NUCLEAR LOCALIZATION SEQUENCE" JOURNAL OF BIOLOGICAL CHEMISTRY, vol. 270, no. 24, 16 June 1995, pages 14255-14258, XP002050723 see the whole document	. 14
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Category '	ation) DOCUMENTS CONSIDERED TO BE RELEVANT  Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
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International application No.

PCT/IB 98/01236

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
Claims Nos.:     because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
Please see additional sheet.
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. X No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:  1-37 all partially
Remark on Protest  The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.

### FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

1. Claims: Invention 1: claims 1-37 all partially

Nucleic acid comprising the sequence as in Seq.ID:38, complementary sequence, fragments, hybridizing sequences. Polypeptide comprising a signal peptide encoded by said nucleotide sequence. Vector encoding a fusion protein comprising said signal peptide. A method of directing the extracellular secretion of a polypeptide by means of said vector. Method of importing a polypeptide into a cell by means of said signal peptide. A method for making a cDNA encoding a secretory protein, partially encoded by said nucleotide sequence, corresponding cDNA. Polypeptide encoded by said nucleotide sequence, comprising a sequence as in Seq.ID:271, method of making said polypeptide. Method of obtaining a promoter located upstream of said nucleotide sequence, promoter thereof.

2. Claims: Invention 2-233: claims 1-37 all partially

Inventions 2-233: Idem as subject 1 but limited to each of the DNA sequences as in Seq.ID:39-271, and corresponding polypeptides, where invention 2 is limited to Seq.ID:39 and 272, invention 3 is limited to Seq.ID:40 and 273,....., invention 233 is limited to Seq.ID:270 and 503).

For the sake of conciseness, the first subject matter is explicitly defined, the other subject matters are defined by analogy thereto.

Ir. nation on patent family members

Internation No PCT/18 98/01236

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